## The Flavor and Fragrance High Production Volume Consortia Revised Robust Summaries for Cinnamyl Derivatives FFHPVC Aromatic Consortium Registration Number

The evaluation of the quality of the following data uses a systematic approach described by Klimisch [Klimisch et al., 1996]. Based on criteria relating to international testing standards for categorizing data reliability, four reliability categories have been established. The following categories are:

• Reliability code 1.

Reliable without restrictions

• Reliability code 2.

Reliable with restrictions

• Reliability code 3.

Not reliable

Reliability code 4.

Not assignable

# OPPT CBIC

#### **Summary of Key Hazard Data for Cinnamyl Derivatives**

ENDPOINT	SUBSTANCE/SURROGATE	VALUE/RANGE <sup>2</sup>	REFERENCE
Vapor pressure	Cinnamaldehyde	0.0289 mm Hg (20°C)	CRC,1973
Vapor pressure	alpha-Amylcinnamaldehyde	0.0012 mm Hg (20°C)	SRC
Vapor pressure	alpha-Hexylcinnamaldehyde	0.0002 mm Hg (20°C)	Vuilleumier, 1995
Vapor pressure	<i>p</i> -t-Butyl- <i>alpha</i> - methylhydrocinnamaldehyde	0.00358 mm Hg (20°C)	SCR
Partition Coefficient	Cinnamaldehyde	1.9	CRC, 1973
Partition Coefficient	alpha-Amylcinnamaldehyde	4.7 (OECD117)	Givaudan, 1994a
Partition Coefficient	alpha-Hexylcinnamaldehyde	5.3 (OECD117)	Givaudan, 1994d
Partition Coefficient	<i>p-t</i> -Butyl- <i>alpha</i> - methylhydrocinnamaldehyde	4.2 (OECD117)	Givaudan, 1994b
Enviloritiental Bate	<b>学等所以</b>	<b>20</b> 40年	作型状态或 <b>产品的图</b> 件指数
Biodegradation3	Cinnamaldehyde	(+) (OECD 301B)	Haarmann& Reimer, 2001
Biodegradation	alpha-Amylcinnamaldehyde	(+) (OECD 301B)	Givaudan, 1992a, Quest, 1996
Biodegradation	alpha-Hexylcinnamaldehyde	(+) (OECD 301B)	Givaudan, 1992b, Quest, 1994

<sup>&</sup>lt;sup>1</sup> Surrogate is a structurally related substance that may include a metabolic product or precursor of the named substance. Range of values may be reported for substance, surrogate or chemical category.

Experimental value or values for a substance or group of substances in the chemical category not biodegradable, (-); readily biodegradable, (+); ready and ultimately biodegradable, (++)

Biodegradation	<i>p-t</i> -Butyl- <i>alpha</i> - methylhydrocinnamaldehyde	(+) (OECD 301F)	Givaudan, 1994c, BBA, 1990
Biodegradation for Category	Cinnamyl Derivatives	Readily Biodegradable	,
Ecotoxicity			
Fish	Cinnamaldehyde	96-hr LC50=4.3 mg/L NOEC=2.8 mg/L	Caspers, 1993
Fish	alpha-Amylcinnamaldehyde	96-hr LC50=3.14 mg/L	SRC
Fish	alpha-Hexylcinnamaldehyde	96-hr LC50=2.36 mg/L	SRC
Fish	<i>p-t</i> -Butyl- <i>alpha</i> - methylhydrocinnamaldehyde	96-hr LC50=3.19 mg/L	SRC
Acute Fish Toxicity Range	Cinnamyl Derivatives	LC50=1-5 mg/L	
Aquatic Invertebrates	Cinnamaldehyde	48-hr EC50=3.86 mg/L, NOEC=1.91 mg/L	Ward, 2003a
		48-hr EC50=11.5 mg/L	Barth &Winkler, 2001
Aquatic Invertebrates	alpha-Amylcinnamaldehyde	48-hr EC50=1.1 mg/L	Caspers, 1993
Aquatic Invertebrate Acute Toxicity	Cinnamyl Derivatives	48-hr EC50=1-5 mg/L	
Aquatic Plant	Cinnamaldehyde	72-hr EC50=4.56 mg/L, NOEC=2.00mg/L (no.cells/ml)	Ward, 2003b
Aquatic Plant	alpha-Amylcinnamaldehyde	72-hr EC50=1.18 mg/L, NOEC=0.154mg/L (no.cells/ml)	Ward, 2003c
Aquatic Plant Acute Toxicity	Cinnamyl Derivatives	72-hr EC50=1-5 mg/L	
Human Health			
Repeat Dose4	Cinnamaldehyde	NOEL=125 mg/kg LOEL=500mg/kg (m&f,r,diet,90d)	Hagan, 1967
		NOEL>200 mg/k (m&f,r,diet,12-wk)	Trubek, 1958b
		NOEL=625mg/kg LOEL=1250mg/kg (m&f,r,diet,90d) NOEL=>200mg/kg (m&f,r,diet,2 yrs)	NTP, 1995
			NTP, 2003
Repeat Dose	alpha-Amylcinnamaldehyde	NOEL=34.9 mg/kg LOEL=320 mg/kg	Carpanini, 1973
Repeat Dose	alpha-Hexylcinnamaldehyde	(m&f,,r,diet,14wk) NOEL=125 mg/kg LOEL=250 mg/kg (m&f,r,dermal,90d)	Lough, 1980
Repeat Dose	<i>p-t</i> -Butyl- <i>alpha</i> -methylhydrocinnamaldehyde	NOEL=25 mg/kg LOEL=50 mg/kg (m,r,dermal,90d)	Givaudan, 1990c,1990d
Repeat Dose Toxicity	Cinnamyl Derivatives	NOEL=25-200 mg/kg	

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<sup>&</sup>lt;sup>4</sup> Value is the NOAEL or NOEL (sex, species route, duration)

		bw/day	
Reproduction	<i>p-t</i> -Butyl- <i>alpha</i> - methylhydrocinnamaldehyde	NOEL=25 mg/kg LOAEL=50 mg/kg (m&f,r,gavage,13wk)	Givaudan, 1990c
		NOEL=25 mg/kg (m&f,r,gavage,13 wk) NOEL>44.6 mg/kg (m&f,d,oral,13wk)	Givaudan, 1990d
		NOEL=25 mg/kg (f,monkey,oral,90d) Pharmacokinetic model	Givaudan, 1990e
		Peak plasma levels and AUC in rats at 25 and 100 mg/kg=100-1000 x plasma levels in humans after maxtopical application	Givaudan, 1990g Hawkins,1994
Reproduction	Cinnamyl alcohol	NOEL>53.5mg/kg (m&f,r,gavage, 11d)	Zaitsev and Maganova, 1975
Reproduction	Cinnamic acid	NOEL>50 mg/kg (m&f,r,gavage, 11d)	Zaitsev and Maganova, 1975
Developmental	Cinnamaldehyde	NOEL>1200 mg/kg (f,m,gavage,d6-15)	Hardin, 1987
In vitro Genotoxicity5	Cinnamaldehyde		Sekizawa and Shibamoto, 1982; Prival et al., 1982; Marnett, 1985; Lijinsky and Andrews, 1980; Kasamaki, 1982; Azizan and Blevins, 1995; Neudecker, 1983
	alpha-Amylcinnamaldehyde	- AMS	Wild, 1983; Fujita and Sasaki, 1987
	alpha-Hexylcinnamaldehyde	- AMS	Wild, 1983
In vivo Genotoxicity	Cinnamaldehyde	-SLR	Woodruff, 1985
	alpha-Amylcinnamaldehyde	-SLR	Wild, 1983
	alpha-Hexylcinnamaldehyde	-SLR	Wild, 1983

<sup>&</sup>lt;sup>5</sup> (-), no significant genotoxic potential; (=/-), equivocal evidence; (+), positive evidence of genotoxicity. AMS, Ames assay; MLA, Mouse Lymphoma assay; ABS, chromosomal aberration assay; UDS, Unscheduled DNA Synthesis; MN, Micronucleus test, SCE, Sister Chromatid Exchange assay, SLA, Sex-linked Lethal assay.

Cinnamaldehyde	-UDS, MN,	Mirsalis,1989; Hayashi, 1984, 1988; Mereto, 1994; NTP, 2003; Sakasi, 1990
alpha-Amylcinnamaldehyde	-MN	Wild, 1983
alpha-Hexylcinnamaldehyde	-MN	Wild, 1983
<i>p-tert</i> -Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde	-MN	Gudi and Krsmanovic, 2000

#### **Robust Summaries**

## 1 Chemical and Physical Properties

## 1.1 Melting Point

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Measured
Remarks for Test Conditions	No test conditions provided.
Melting Point	-7.5
Remarks for Results	The data are considered reliable.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
References	Merck & Co., Inc. The Merck Index (1997) 12th Edition, Publishers: Merck Research Laboratories, Whitehouse Station, NJ.
Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Measured
Remarks for Test Conditions	No test conditions provided.
Melting Point	80 °C
Remarks for Results	The data are considered reliable.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
References	Handbook of Chemistry and Physics (CRC) 54 <sup>th</sup> Edition (1973) Publishers: CRC Press, Cleveland, Ohio.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Mean or weighted
Melting Point	0.04 °C
Remarks for Data	Calculated

References	Syracuse Research (	Corporation (SRC)
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Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Mean or weighted
Melting Point	33.9 °C
Remarks for Data	Calculated
References	Syracuse Research Corporation (SRC)
Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Mean or weighted
Melting Point	44.4 °C
Remarks for Data	Calculated
References	Syracuse Research Corporation (SRC)
Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	Mean or weighted
Melting Point	46.3 °C
Remarks for Data	Calculated
References	Syracuse Research Corporation (SRC)
Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Melting Point	4 °C
References	Fenaroli's Handbook of Flavor Ingredients Volume II 3rd Edition. Edited by G. Burdock. CRC Press, 1994, Reston VA.

## 1.2 Boiling Point

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
<b>Boiling Point</b>	246 °C
Remarks for Test Conditions	No test conditions provided
References	Handbook of Chemistry and Physics (CRC) 54 <sup>th</sup> Edition (1973) Publishers: CRC Press, Cleveland, Ohio.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Boiling Point	250 °C
Remarks for Test Conditions	No test conditions provided
References	Fragrance Materials Association (FMA)

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
<b>Boiling Point</b>	284 °C
Remarks for Test Conditions	No test conditions provided
References	Fragrance Materials Association (FMA)

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
<b>Boiling Point</b>	226.7 °C
Method/guideline	Stein and Brown Method
Remarks for Test Conditions	Calculated
References	Syracuse Research Corporation (SRC)

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7

**Boiling Point** 304.8 °C

Method/guideline Stein and Brown Method

Remarks for Test Conditions Calculated

**References** Syracuse Research Corporation (SRC)

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Boiling Point	305 °C
Remarks for Test Conditions	No test conditions provided

**References** Fragrance Materials Association (FMA)

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
<b>Boiling Point</b>	318.7 °C
Method/guideline	Stein and Brown Method
Remarks for Test Conditions	Calculated

**References** Syracuse Research Corporation (SRC)

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Boiling Point	280 °C
Method/guideline	Stein and Brown Method
Remarks for Test Conditions	Calculated
References	Syracuse Research Corporation (SRC)

Substance Name	p-t-Butyl-alpha-methylhydrocinnamaldehyde
CAS No.	80-54-6
<b>Boiling Point</b>	258°C
References	Arctander's Perfume and Flavor Chemicals Vol. I Publisher: S. Arctander (1969) Montclair, NJ

#### 1.3 Vapor Pressure

Cinnamaldehyde
104-55-2
Measured
NA
0.0289mm Hg (0.00347 kPa)
20 °C
Handbook of Chemistry and Physics (CRC) 54 <sup>th</sup> Edition (1973) Publishers: CRC Press, Cleveland, Ohio.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Calculated
GLP	NA
Vapor Pressure	0.02mm Hg (0.00267 kPa)
Temperature	20 °C
References	Fragrance Materials Association (FMA)

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Modified Antoine and Grain Method
GLP	NA
Vapor Pressure	0.09 mm Hg (0.012 kPa)
Temperature	20 °C
References	Syracuse Research Corporation (SRC)

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Calculated

**GLP** NA

**Vapor Pressure** <0.001 mm Hg (<0.00013 kPa)

Temperature 20 °C

**References** Fragrance Materials Association (FMA)

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Modified Grain Method
GLP	NA
Vapor Pressure	0.0012 mm Hg ((0.00016 kPa)
Temperature	20 °C
References	Syracuse Research Corporation (SRC)

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
GLP	NA
Year	1995
Vapor Pressure	0.0002 mg Hg (0.000027 kPa)
Temperature	20 °C
References	Vuilleumier C., Flament, I., Sauvegrain, P. (1995) Headspace measurement of evaporation rates of perfumes applied onto the skin: Application to rose essential oils and their principal components. Perfumer and Flavorish 20(2), 1-9.

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Calculated
GLP	NA
Vapor Pressure	<0.001 mm Hg (<0.00013 kPa)
Temperature	20 °C
References	Fragrance Materials Association (FMA)

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Modified Grain Method
GLP	NA
Vapor Pressure	0.00048 mm Hg (0.000064 kPa)
Temperature	20 °C
References	Syracuse Research Corporation (SRC)

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde	
CAS No.	80-54-6	
Method/guideline	Modified Grain Method	
GLP	NA	
Vapor Pressure	0.00358 mm Hg (0.00048 kPa)	
Temperature	20 °C	
References	Syracuse Research Corporation (SRC)	

#### 1.4 Octanol/Water Partition Coefficient

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
GLP	NG
Year	NG
Method/guideline	Measured
Partition coefficient	1.9
Remarks for Data Reliability	Comparable to guidelines/standards.
Data Quality Reliabilities	Reliability code 2. Reliable with restrictions.
References	Syracuse Research Corporation (SRC)-(Hansch et al, 1995)

Substance Name	alpha-Hexylcinnamaldehyde	
CAS No.	101-86-0	
Method/guideline	OECD Guideline No. 117	
GLP	Yes	
Year	1994	
Log Pow	5.3	
Temperature	24 °C	
Remarks for Data Reliability	Guideline study. The log Kow compares well with the calculated value. Data are considered reliable.	
Data Quality Reliabilities	Reliability 1. Reliable without restriction.	
References	Givaudan-Roure (1994d) Partition coefficient n-octanol/water of alpha-hexylcinnamaldehyde. Unpublished report to RIFM.	

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
GLP	NG
Year	NG
Method/guideline	Calculated
Partition coefficient	1.82
Remarks for Data Reliability	Comparable to guidelines/standards.
Data Quality Reliabilities	Reliability code 2. Reliable with restrictions.
References	Syracuse Research Corporation (SRC)

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
GLP	Yes
Year	1994
Method/guideline	OECD Guideline No. 117
Log Pow	4.7
Temperature	24 °C
Remarks for Data Reliability	Guideline study. The log Kow compares well with the calculated value. Data are considered reliable.

**Data Quality Reliabilities** Reliability code 1. Reliable without restriction.

**References** Givaudan-Roure (1994a) Partition coefficient n-octanol/water of

alpha-amylcinnamaldehyde. Unpublished report to RIFM.

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
GLP	NG
Year	NG
Method/guideline	Calculated
Log Pow	4.33
Remarks for Data Reliability	Comparable to guidelines/standards.

Substance Name	alpha-Hexylcinnamaldehyde	

Reliability code 2. Reliable with restrictions.

Syracuse Research Corporation (SRC)

**CAS No.** 101-86-0

GLP NG Year NG

**Data Quality Reliabilities** 

References

Method/guideline Calculated

**Log Pow** 4.82

**Remarks for Data Reliability** Comparable to guidelines/standards.

**Data Quality Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Syracuse Research Corporation (SRC)

Substance Name	<i>alpha</i> -Hexylcinnamaldehyde
CAS No.	101-86-0
GLP	NG
Year	1996
Method/guideline	Measured
Log Pow	4.9

Remarks for Data Reliability Comparable to guidelines/standards.

**Data Quality Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Quest (1994) Private communication to FMA.

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde	
CAS No.	80-54-6	
GLP	NG	
Year	NG	
Method/guideline	Calculated	
Log Pow	4.36	
Remarks for Data Reliability	Comparable to guidelines/standards.	
Data Quality Reliabilities	Reliability code 2. Reliable with restrictions.	
References	Syracuse Research Corporation (SRC)	

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde	
CAS No.	80-54-6	
GLP	Yes	
Year	1994	
Method/guideline	OECD Guideline No. 117	
Log Pow	4.2	
Temperature	24 °C	
Remarks for Data Reliability	Guideline study. The log Kow compares well with the calculated value. Data are considered reliable.	
Data Quality Reliabilities	Reliability 1. Reliable without restriction.	
References	Givaudan-Roure (1994b) Partition coefficient n-octanol/water of <i>p</i> -t-butyl-alpha-methyldihydrocinnamic aldehyde. Unpublished Report to RIFM.	

#### 1.5 Water Solubility

Substance Name	Cinnamaldehyde	
CAS No	104-55-2	
LASNO	1114-55-7	

Method/guideline	Measured
Value (mg/L) at temperature	1420 mg/L (20 °C)
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	Syracuse Research Database

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Calculated at log Kow=1.90 (ESPKOW)
Value (mg/L) at temperature	2150 mg/L
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	ESPOW

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Calculated at log Kow=4.33 (ESPKOW)
Value (mg/L) at temperature	8.5 mg/L
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	ESPKOW

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Calculated at log Kow=4.82 (ESPKOW)
Value (mg/L) at temperature	2.75 mg/L
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	ESPKOW

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	80-54-6
Year	1995

Method/guideline **OECD 105** 

Value (mg/L) at temperature 33 mg/L at 20 °C

Comparable to guidelines/standards. Reliability code 1. Reliable without restrictions. **Remarks for Data Reliability** 

References Givaudan-Roure (1995) Water solubility of p-t-butyl-alpha-

methylhydrocinnamic aldehyde. Unpublished Report to RIFM.

methylhydrocinnamic aldehyde. Unpublished Report to RIFM.

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	80-54-6
Year	1994
Method/guideline	NG
Value (mg/L) at temperature	<100 mg/L at 20 °C
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	Givaudan-Roure (1995) Water solubility of <i>p</i> -t-butyl-alpha-

Substance Name	<i>p</i> -t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	80-54-6
Year	1990
Method/guideline	NG
Value (mg/L) at temperature	0.02% w/v (200 mg/L)
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	Bush Boake Allen (BBA) (1990). Biodegradability of p-t-butylalpha-methylhydrocinnamic aldehyde and methyl-alpha-ionone. Unpublished report to RIFM.

Substance Name	<i>p-</i> t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	Calculated at log Kow= 4.36 (ESKOW)
Value (mg/L) at temperature	7.8 mg/L
Remarks for Data Reliability	Comparable to guidelines/standards. Reliability code 2. Reliable with restrictions.
References	ESPKOW

## 2 Environmental Fate and Pathways

## 2.1 Photodegradation

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Calculation
Test Type	AOPWIN
Halflife t1/2	3.17
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.
References	AOPWIN

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Method/guideline	Calculation
Test Type	AOPWIN
Halflife t1/2	2.40 hrs
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.
References	AOPWIN

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Calculation
Test Type	AOPWIN
Halflife t1/2	2.33 hrs
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.
References	AOPWIN

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	Calculation
Test Type	AOPWIN
Halflife t1/2	3.88 hrs
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.
References	AOPWIN

## 2.2 Biodegradation

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Biodegradability was determined by sealed vessel test based on OECD Guideline 301 B.
Test Type	OECD 301 B CO2 evolution
GLP	Yes
Year	1996
Contact Time (units)	28 days
Innoculum	Secondary effluent from an unacclimatized activate
Degradation % after time	89% at 7 days, 94% at 14 days and 100% at 21, 27, and 28 days
Time required for 10% degradation	1 day
10 day window criteria	Yes
Total degradation	100%
Conclusion Remarks	Readily biodegradable
Data Qualities Reliabilities	Reliability code 1. Reliable without restriction.
Remarks for Data Reliability	The study was conducted in accordance with OECD 301B guidelines.
References	Haarmann and Reimer (2001) Ready Biodegradability of Cinnamic Aldehyde according to OECD Guideline No. 301B Private Communication to RIFM.
Substance Name	p-t-Butyl-alpha-methyldihydrocinnamic aldehyde,
Substance Name	91-98% pure, clear, almost colorless liquid, fresh, light, green

floral, reminiscent of lily; strongly diffusive

**CAS No.** 80-54-6

Method/guideline OECD Guideline 301F

Test Type Respirometric Method, MITI Test I

GLP Yes

**Year** 1994

Contact Time (units) 28 days

**Innoculum** Acitivated sludge fro a local sewage plant

Remarks for Test Conditions Samples were incubated with acitivated sludge for 28 days.

Reference material was 100 mg/L aniline. BOD was measured

**Degradation % after time** 68% at 28 days

Results At 50 mg/L, 84% biodegradation was recorded at 28 days and

78% in 10 day window (days 8 to 18).

At 100 mg/L, 68% biodegradation was recorded at 28 days and

57% in 10 day window (days 8 to 18)

Time required for 10%

degradation

10 day window criteria

NA

Yes

Total degradation NA

**Conclusion Remarks**The shape of the biodegradation curve indicated the

occurrence of an adaptation period for biodegradation.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

Remarks for Data Reliability The study was conducted in accordance with OECD 301F

quidelines.

**References** Givandan-Roure (1994c) Ready biodegradability of *p*-t-butyl-

alpha-methylhydrocinnamic aldehyde Unpublished report to

RIFM.

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamic aldehyde,
	91-98% pure, clear, almost colorless liquid, fresh, light, green
	floral, reminiscent of lily; strongly diffusive

**CAS No.** 80-54-6

Method/guideline OECD Guideline 111 & Directive 92/69/EEC, Method C.7

**Test Type** Abiotic degradation

GLP NG

**Year** 1995

Contact Time (units) 16 days

**Innoculum** None; Aqueous solution (10 mg/L) in air

**Remarks for Test Conditions** Samples were agitated in air, regularly removed and analyzed

for the test substances by HPLC.

Degradation % after time >50% at 16 days

Results Oxidation reactions follow pseudo first order kinetics.

Time required for 10%

degradation

NA

10 day window criteria NA

Total degradation NA

**Conclusion Remarks** The authors concluded that the test substance has an

> approximately lifetime of two weeks in the environment. The corresponding acid is the major oxidation product identified.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

The study was conducted in accordance with OECD 111 Remarks for Data Reliability

guidelines.

Givandan-Roure (1995b) Stability of p-t-butyl-alpha-References

methylhydrocinnamic aldehyde in aqueous solution in the

presence of air. Unpublished report to RIFM.

p-t-Butyl-alpha-methyldihydrocinnamic aldehyde, **Substance Name** 

91-98% pure, clear, almost colorless liquid, fresh, light, green

floral, reminiscent of lily; strongly diffusive

CAS No. 80-54-6

Method/guideline Method F

**Test Type** DOC - Method F from Blue book series, 1981

**GLP** NG

1990 Year

**Contact Time (units)** 28 days

Innoculum Activated sludge from local STP

**Remarks for Test Conditions** 50.04 mg DOC/L at 20 C for 28 days

Degradation % after time 96% at 31 days

Results 92 % biodegradation after 28 days. 96% after day 31.

Time required for 10%

degradation

<1 day

10 day window criteria

Yes

Total degradation Yes

Conclusion Remarks Readily biodegradable

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction. guidelines.

**References** Bush Boake Allen (BBA) (1990). Biodegradability of *p*-t-butyl-

alpha-methylhydrocinnamic aldehyde and methyl-alpha-ionone.

Unpublished report to RIFM.

Substance Name alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Method/guideline Biodegradability was determined by sealed vessel test based

on OECD Guideline 301 B.

**Test Type** OECD 301 B CO2 evolution

**GLP** Yes

**Year** 1996

Contact Time (units) 28 days

Innoculum Secondary effluent from an unacclimatized activate

**Degradation % after time** 65% at 28 days

Time required for 10%

degradation

9 days

10 day window criteria Yes

Total degradation No

Conclusion Remarks Readily biodegradable

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

auidelines.

**References** Givaudan-Roure (1989) Ready Biodegradability of Amyl

Cinnamic Aldehyde according to OECD Guideline No. 301B

Private Communication to FMA.

Substance Name alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Method/guideline Ready biodegradability of amyl acinnamic aldehyde was

determined according to OECD Guideline No. 301F.

Test Type OECD No. 301F Respirometric method/ SAPROMAT

**GLP** Yes

**Year** 1992

Contact Time (units) 28 days

**Innoculum** Activated sludge

Remarks for Test Conditions Bottle 1 & 2: Basal culture medium + activated sludge 30 mg/l +

test chemical (100 mg/l); Bottle 3: Basal culture medium + activated sludge 30mg/l+aniline (100mg/l); Bottle 4: Basal

culture medium + activated sludge 30 mg/l.

**Degradation % after time** 90% in 28days

**Results** 90% of the test chemical was biodegraded in 28 day as

compared to only 61% of reference material (aniline) was

biodegraded in 28 days.

Total degradation Yes

Conclusion Remarks Readily biodegradable

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

Remarks for Data Reliability The study was conducted in accordance with OECD 301F

guidelines.

**References** Givaudan Roure (1992a). Ready Biodegradability of Amyl

Cinnamic Aldehyde according to OECD Guideline No. 301F.

Unpublished report to RIFM.

Substance Name alpha-Hexylcinnamaldehyde: Pale yellow oily liquid with sweet

slightly floral odor.

**CAS No.** 101-86-0

Method/guideline Sealed vessel test: based on OECD Guideline 301B

Test Type OECD 301B CO2 evolution

**GLP** Yes

**Year** 1994

Contact Time (units) 28 days

Innoculum Secondary effluent from unacclimatized activated sludge plant

Remarks for Test Conditions Test concentration: 11.9 mg/l organic carbon. Test temp: 20-24

°C

**Degradation % after time** 76.5% at 28 days

**Results** 76.5% biodegradable (95% CI-67.0-85.9) in 28 days.

Time required for 10%

degradation

<11 days

10 day window criteria No

Total degradation No

Conclusion Remarks The test substance achieved the 60% pass level by day 28 but

failed the 10 day window criterium and therefore can be classified as ultimately biodegradable according to this test

protocol.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

guidelines.

Quest (1994) Report on Hexyl Cinnamic Aldehyde Biodegradation. References

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Ready Biodegradability of the test material was determined according to OECD Guideline No. 301F
Test Type	OECD No. 301F, Respirometric method
GLP	Yes
Year	1992
Contact Time (units)	28 days
Innoculum	Activated sludge
Remarks for Test Conditions  Degradation % after time	Bottle 1 & 2: Basal culture medium + activated sludge 30 mg/l + test chemical (~100 mg/l). Bottle 3: Basal culture medium + activated sludge 30 mg/l + aniline (~100 mg/l); Bottle 4: Basal culture medium + activated sludge 30 mg/l. 97% in 28 days
Results	97% of the test material was biodegraded in 28 days as compare to 61% of aniline in the same period.
Total degradation	Yes
<b>Conclusion Remarks</b>	Readily biodegradable
Data Qualities Reliabilities	Reliability code 1. Reliable without restriction.
Remarks for Data Reliability	The study was conducted in accordance with OECD 301F guidelines.
References	Givaudan Roure. (1992b). Ready Biodegradability of Hexyl Cinnamic Aldehyde according to OECD Guideline No. 301F. Unpublished report to RIFM.

Substance Name	alpha-Hexylcinnamaldehyde 94% pure - 44% cis and 50% trans
CAS No.	5392-40-5
Test Type	OECD 301B CO2 evolution
GLP	No
Year	1994
Contact Time (units)	28 days
Innoculum	Secondary effluent from sludge from local STP
Remarks for Test Conditions	10 mg/l organic carbon at 20 °C for 28 days

**Degradation % after time** 92.1% at 28 days

**Results** 92.1% biodegradation in 28 days

Time required for 10%

degradation

<4 days

10 day window criteria Yes

Total degradation No

**Conclusion Remarks** Readily biodegradable

**Data Qualities Reliabilities** Reliability code 1. Reliable without restriction.

guidelines.

**References** Quest (1994) Private communication to FMA.

#### 2.3 Fugacity

Substance	Cinnamaldehyde
CAS	104-55-2
Model Conditions	25 C, 1000 lbs.
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay
Model Used (title, version, date)	EQC Fugacity Level III
Input parameters	MW, VP(0.0289 mm Hg), log Kow (1.9), water solubility (1420 mg/L), MP, -7.5 °C, BP, 246.°C
Year	2000
Media	Air-Water-Soil-Sediment Partition Coefficients
Model data and results	Compartment half-lives, hours: Air=7.77; Water=900;Soil=900;Sediment=3600
Estimated Distribution and	Air=0.597%
Media Concentration	Water=25.6%
	Soil=68.4%
	Sediment=5.47%
Conclusion remarks	Substance is predicted to persist in the environment for 632
	hours.
Reliabilities	Reliability code 4. Not assignable.
Remarks for Data Reliability	The data are obtained by a recognized fugacity calculation
	method. Data are considered
	reliable with restriction because this method does not allow
	for biodegradation or

#### metabolism.

References	Mackay D., A.DiGuardo, S.Paterson and C.E.Cowan (1996b) Evaluating the fate of a variety of types of chemicals using the EQC model. Environmental Toxicology and Chemistry, 15(9), 1627-1637.
Substance	alpha-Amylcinnamaldehyde
CAS	122-40-7
Model Conditions	25 C, 1000 lbs.
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay
Model Used (title, version, date)	EQC Fugacity Level III
Input parameters	MW, VP(0.0012 mm Hg), log Kow (4.7), water solubility (8.5 mg/L), MP, 80 °C, BP, 284.°C
Year	2000
Media	Air-Water-Soil-Sediment Partition Coefficients
Model data and results	Compartment half-lives, hours: Air=3.47; Water=360;Soil=360;Sediment=1400
Estimated Distribution and Media Concentration	Air=0.575% Water=32.6% Soil=57.3% Sediment=9.53%
Conclusion remarks	Substance is predicted to persist in the environment for 302 hours.
Reliabilities	Reliability code 4. Not assignable.
Remarks for Data Reliability	The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.
References	Mackay D., A.DiGuardo, S.Paterson and C.E.Cowan (1996b) Evaluating the fate of a variety of types of chemicals using the EQC model. Environmental Toxicology and Chemistry, 15(9), 1627-1637.
Substance	alpha-Hexylcinnamaldehyde
CAS	101-86-0
Model Conditions	25 C, 1000 lbs.
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay

Model Used (title, version, date)	EQC Fugacity Level III
Input parameters	MW, VP(0.0002 mm Hg), log Kow (5.3), water solubility (2.75 mg/L), MP, 44 °C, BP, 305.°C
Year	2000
Media	Air-Water-Soil-Sediment Partition Coefficients
Model data and results	Compartment half-lives, hours: Air=3.41; Water=360;Soil=360;Sediment=1400
Estimated Distribution and Media Concentration	Air=0.449% Water=25.4% Soil=47.7% Sediment=26.5%
Conclusion remarks	Substance is predicted to persist in the environment for 364 hours.
Reliabilities	Reliability code 4. Not assignable.
Remarks for Data Reliability	The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.
References	Mackay D., A.DiGuardo, S.Paterson and C.E.Cowan (1996b) Evaluating the fate of a variety of types of chemicals using the EQC model. Environmental Toxicology and Chemistry, 15(9), 1627-1637.
Substance	<i>p-</i> t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS	80-54-6
Model Conditions	25 C, 1000 lbs.
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay
Model Used (title, version, date)	EQC Fugacity Level III
Input parameters	MW, VP(0.00358 mm Hg), log Kow (4.2), water solubility (33 mg/L), MP, 46 °C, BP, 258.°C
Year	2000
Media	Air-Water-Soil-Sediment Partition Coefficients
Model data and results	Compartment half-lives, hours: Air=7.77; Water=900;Soil=900;Sediment=3600
Estimated Distribution and Media Concentration	Air=0.597% Water=25.6% Soil=68.4% Sediment=5.47%
Conclusion remarks	Substance is predicted to persist in the environment for 632 hours.
Reliabilities	Reliability code 4. Not assignable.

Remarks for Data Reliability	The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.
References	Mackay D., A.DiGuardo, S.Paterson and C.E.Cowan (1996b) Evaluating the fate of a variety of types of chemicals using the EQC model. Environmental Toxicology and Chemistry, 15(9), 1627-1637.

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Model Conditions	25 °C, 100,000 lbs
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay
Model Used	EQC V 2.11 Level 1
Input Parameters	MW, log Kow, water solubility, VP, estimated MP
Media	Air-Water Partition Coefficient
Absorption Coefficient	0.0099
Data Qualities Reliabilities	Reliable with restriction
Remarks for Data Reliability References	The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning
Neierences	Model Version 2.11. Based on Mackay, Donald (1991)  Multimedia environmental models: The fugacity approach.  Lewis Publications, CRC Press, Boca Raton, FL

Substance Name	alpha-amylcinnamaldehyde
CAS No.	122-40-7
Model Conditions	25 °C, 100,000 lbs
Test Type	Environmental Equilibrium Partitioning Model
Method	Mackay
Model Used	EQC V 2.11 Level 1
Input Parameters	MW, log Kow, water solubility, VP, estimated MP
Media	Soil-Water Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

> method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** *alpha-*Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Sediment-Water Partition Coefficient

**Absorption Coefficient** 1970

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** *alpha*-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, VP, estimated MP

Media Suspended Sediment-Water Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** *alpha*-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Fish-Water Partition Coefficient

Absorption Coefficient 2510

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Aerosol-Air Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

**Remarks for Data Reliability** The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Air

**Estimated Distribution and Media Concentration** 

9.7%

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

References Level 1 Fugacity-based Environmental Equilibrium Partition Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** *alpha*-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Water

Estimated Distribution and Media Concentration

1.94%

Data Qualities Reliabilities

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Multimedia environmental models: The fugacity appro

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Soil

Estimated Distribution and Media Concentration

86.4%

Data Qualities Reliabilities

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Model Conditions 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Sediment

**Estimated Distribution and Media Concentration** 

1.92%

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

122-40-7 CAS No.

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Suspended Sediment

**Estimated Distribution and** Media Concentration

0.06%

Data Qualities Reliabilities

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

CAS No. 122-40-7

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, VP, estimated MP **Input Parameters** 

Media Fish **Estimated Distribution and Media Concentration** Data Qualities Reliabilities

Remarks for Data Reliability

References

Reliable with restriction

0.0049%

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Amylcinnamaldehyde

122-40-7 CAS No.

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, VP, estimated MP

Media Aerosol

**Estimated Distribution and** Media Concentration

Remarks for Data Reliability

0.0018%

Data Qualities Reliabilities Reliable with restriction

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

The data are obtained by a recognized fugacity calculation

References Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

CAS No. 101-86-0

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Air-Water Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Soil-Water Partition Coefficient

Absorption Coefficient 3230

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

Model Conditions 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Sediment-Water Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** *alpha*-hexylcinnamaldehyde

**CAS No.** 101-86-0

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Suspended Sediment-Water Partition Coefficient

Absorption Coefficient 24500

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Fish-Water Partition Coefficient

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Aerosol-Air Partition Coefficient

**Absorption Coefficient** 14100000

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

**Media** Air

**Estimated Distribution and Media Concentration** Data Qualities Reliabilities

5.7%

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

101-86-0 CAS No.

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Water

**Estimated Distribution and Media Concentration** 

0.52%

Data Qualities Reliabilities

References

Reliable with restriction

Remarks for Data Reliability

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

CAS No. 101-86-0

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Soil

**Estimated Distribution and Media Concentration** Data Qualities Reliabilities

91.7%

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

101-86-0 CAS No.

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Sediment

**Estimated Distribution and** Media Concentration

2.0%

Data Qualities Reliabilities

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

CAS No. 101-86-0

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Suspended Sediment

**Estimated Distribution and Media Concentration** 

Data Qualities Reliabilities

0.064%

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

101-86-0 CAS No.

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Fish

**Estimated Distribution and** Media Concentration

0.0052%

Data Qualities Reliabilities

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** alpha-Hexylcinnamaldehyde

CAS No. 101-86-0

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Aerosol Estimated Distribution and Media Concentration Data Qualities Reliabilities

0.0016%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Air-Water Partition Coefficient

Absorption Coefficient 0.0031

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References

Level 1 Fugacity-based Environmental Equilibrium Partitioning
Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Soil-Water Partition Coefficient

**Absorption Coefficient** 1.30

References

Data Qualities Reliabilities Reliable with restriction

> method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Sediment-Water Partition Coefficient

Absorption Coefficient 2.60

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Suspended Sediment-Water Partition Coefficient

**Absorption Coefficient** 8.12

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Fish-Water Partition Coefficient

Absorption Coefficient 3.30

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Aerosol-Air Partition Coefficient

**Absorption Coefficient** 483000

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Air

**Estimated Distribution and Media Concentration** 

12.7%

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

References Level 1 Fugacity-based Environmental Equilibrium Partitic Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Water

Estimated Distribution and Media Concentration Data Qualities Reliabilities

82.4%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Soil

Estimated Distribution and Media Concentration Data Qualities Reliabilities

4.82%

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Sediment

**Estimated Distribution and Media Concentration** Data Qualities Reliabilities

0.11%

Reliable with restriction

**Remarks for Data Reliability** 

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** Cinnamaldehyde

CAS No. 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Suspended Sediment Media

**Estimated Distribution and** Media Concentration

0.0034%

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach. Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** Cinnamaldehyde

CAS No. 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Fish **Estimated Distribution and Media Concentration** Data Qualities Reliabilities

0.00027%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** Cinnamaldehyde CAS No. 104-55-2

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Aerosol

**Estimated Distribution and** Media Concentration Data Qualities Reliabilities

0.00012%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Air-Water Partition Coefficient Absorption Coefficient 0.001

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Soil-Water Partition Coefficient

**Absorption Coefficient** 312

Data Qualities Reliabilities Reliable with restriction

**Remarks for Data Reliability** The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Sediment-Water Partition Coefficient

**Absorption Coefficient** 624

References

Data Qualities Reliabilities Reliable with restriction

**Remarks for Data Reliability** The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Suspended Sediment-Water Partition Coefficient

**Absorption Coefficient** 1950

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Fish-Water Partition Coefficient

**Absorption Coefficient** 792

References

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991)
Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Aerosol-Air Partition Coefficient

**Absorption Coefficient** 15000000

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism.

References Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

**Media** Air

**Estimated Distribution and Media Concentration** Data Qualities Reliabilities

3.2%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Water

**Estimated Distribution and** Media Concentration

6.3%

Data Qualities Reliabilities

Reliable with restriction

Remarks for Data Reliability

References

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

The data are obtained by a recognized fugacity calculation

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Soil Estimated Distribution and Media Concentration Data Qualities Reliabilities

88.5%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Sediment

Estimated Distribution and Media Concentration

References

2.0%

Data Qualities Reliabilities Reliable with restriction

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

Model Conditions 25 °C, 100,000 lbs

Test Type Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

Input Parameters MW, log Kow, water solubility, MP, estimated VP

Media Suspended Sediment

**Estimated Distribution and Media Concentration** Data Qualities Reliabilities

0.061%

Reliable with restriction

Remarks for Data Reliability

References

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

**Model Conditions** 25 °C, 100,000 lbs

**Test Type** Environmental Equilibrium Partitioning Model

Method Mackay

Model Used EQC V 2.11 Level 1

**Input Parameters** MW, log Kow, water solubility, MP, estimated VP

Media Fish

**Estimated Distribution and** Media Concentration

References

0.0050%

Reliable with restriction Data Qualities Reliabilities

Remarks for Data Reliability The data are obtained by a recognized fugacity calculation

> method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning

Model Version 2.11. Based on Mackay, Donald (1991)

Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

**Substance Name** p-t-butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

25 °C, 100,000 lbs **Model Conditions** 

**Test Type Environmental Equilibrium Partitioning Model** 

Method Mackay

Model Used EQC V 2.11 Level 1

MW, log Kow, water solubility, MP, estimated VP **Input Parameters** 

Media Aerosol Estimated Distribution and Media Concentration Data Qualities Reliabilities

0.0010%

Reliable with restriction

Remarks for Data Reliability

The data are obtained by a recognized fugacity calculation method. Data are considered reliable with restriction because this method does not allow for biodegradation or metabolism. Level 1 Fugacity-based Environmental Equilibrium Partitioning Model Version 2.11. Based on Mackay, Donald (1991) Multimedia environmental models: The fugacity approach.

Lewis Publications, CRC Press, Boca Raton, FL

# 3 Ecotoxicity

References

#### 3.1 Acute Toxicity to Fish

Substance Name	Cinnamaldehyde
CAS Numerical	106-24-1
Remarks for Substance	Assay: >99%
Method/guideline	96-Hour semi-static toxicity test
GLP	Yes (Council Directive 92/69/EEC C.1 (1992)
Year	1993
Test Type	Experimental
Species/Strain/Supplier	Zebra fish/ <i>Brachydanio rerio</i> /Hamilton Buchanan/West Aquarium
Remarks for Test Conditions	Zebra fish (6-month old, length, 2.5-3.5 cm) were exposed to a series of 5 test concentrations of 0, 2.8, 3.9, 5.5, and 7.8 mg/L of cinnamaldehyde for 96 hours in a semi-static test. Ultra turrax was used as a solubilizer. Solutions were renewed every 24 hours. Fish were maintained on a schedule of 12 hours of light and 12 hours of darkness. Fish were observed twice daily for mortality and symptoms. GC analytical measurement of cinnamaldehyde concentrations in control solutions, <1 mg/L.
Conclusion Remarks	The acute 96-hour LC50 for cinnamaldehyde in zebra fish is 3.1 mg/L
Remarks for Results	Measured concentrations at 0 and 24 hours were: <0.25 mg/L for controls 2.63 and 2.68 mg/L at 0 hours and 0.87 and 1.94 mg/L at 24 hours for nominal concentration of 2.8 mg/L 3.75 and 3.69 mg/L at 0 hours and 2.12 and 2.62 mg/L at 24 hours for nominal concentration of 3.9 mg/L 5.42 and 5.23 mg/L at 0 hours and 3.67 and 4.18 mg/L at 24 hours for nominal concentration of 5.5 mg/L 7.43 and 7.09 mg/L at 0 hours and 5.54 and 6.72 mg/L at 24 hours for nominal concentration of 7.8 mg/L

Temperature, oxygen, oxygen saturation and pH measured at 0 and 24 hours were:

20.8-22.6 C, 8.3-9.1 mg/L, 99.9-110%, and 7.4-8.2 for controls; 20.9-22.7 C, 8.2-9.5 mg/L, 95.6-109.2%, and 7.5-8.2 at 2.8 mg/L.

21.1-22.7 C, 8.4-9.6 mg/L, 97.9-113%, and 7.5-8.2 at 3.9 mg/L 21.2-22.7 C, 8.6-9.6 mg/L, 99.9-111.5%, and 7.7-8.2 at 5.5 mg/L.

21.1-22.8 C, 8.6-9.5 mg/L, 99.9-113%, and 7.6-8.2 at 7.8 mg/L.

The cumulative number of deaths at each 24- hour intervals and the corresponding concentrations in mg/L at t=0, 24,48, 72, & 96 hours are:

control and 2.8 mg/L, 0/10 at all times;

at 3.9 mg/L, 0/10 at 0, 24, and 48 hours, 2/10 at 96 hours; at 5.5 mg/L, 0/10 at 0, 24, and 48 hours, 9/10 at 72 hours and

10/10 at 96 hours

at 7.8 mg/L, 1/10 at 24 hours, and 10/10 at 48 hours

Analytical monitoring GC

Unit mg/L

Exposure period (unit) 96 hours

Nominal concentrations as

mg/L

Measured concentrations as

mg/L

**Endpoint value** 96-hour LC0=2.8, 96-hour LC50=4.3 (plot of log conc vs %

0, 2.8, 3.9, 5.5, and 7.8 mg/L

mortality), and 96-hour LC100=5.5 mg/L

**Data Qualities Reliabilities** Reliability code 2. Reliable with restriction.

Remarks for Data Reliability Code 2. Acceptable, well-documented publication/study report

which meets basic scientific principles.

**References** Caspers (1993) Acute fish toxicity of cinnamic aldehyde. Study

No. 341 A/92 F. Private Communication to RIFM. Unpublished

report.

Substance Name	Cinnamic aldehyde
CAS No.	104-55-2
Method/guideline	ECOSAR
Test Type	Calculated based on log Kow
Species/Strain/Supplier	Fish
Exposure period	96 hr
Conclusion Remarks	LC50 = 11.9 mg/l

### Remarks for Data Reliability

The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.

	alpha-Amylcinnamaldehyde
Substance Name	
CAS No.	122-40-7
Method/guideline	ECOSAR
Test Type	Calculated based on measured Kow
Species/Strain/Supplier	Fish
Exposure period	96 hr
Conclusion Remarks	LC50 = 3.14 mg/l
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method but are not consistent with chemical structure. Data are considered overly conservative.
Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	ECOSAR
Test Type	Calculated based on measured Kow
Species/Strain/Supplier	Fish
Exposure period	96 hr
<b>Conclusion Remarks</b>	LC50 = 2.36 mg/l
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method but are not consistent with chemical structure. Data are considered overly conservative.
Substance Name	p-tert-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	ECOSAR
Test Type	Calculated based on measured Kow
Species/Strain/Supplier	Fish
Exposure period	96 hr
<b>Conclusion Remarks</b>	LC50 = 3.19  mg/l
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation

# 3.2 Acute Toxicity to Aquatic Invertebrates

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7
Remarks for Substance	Assay: 99%
Method/guideline	Acute toxicity for Daphnia, Directive 67/548/EWG (1992)
Test Type	Experimental (48- hr static test)
GLP	Yes
Year	1992
Analytical procedures	GLC
Species/Strain	Daphnia magna/Strauss-pathogen free.
Test details	48 hrs.
Remarks for Test Conditions	Parthenogenetic female daphnids (6-24 hours old) produced from a laboratory culture of adults. 40 daphnid were randomly selected for each replicate test. Tests were performed at 5 nominal concentrations. Mortality, immobility, and sub-lethal effects were determined visually at 0, 24, and 48 hours. Test temperature was maintained at 20.3-20.5 C
Nominal concentrations as mg/L	0, 0.2, 0.4, 0.7, 1.4, 2.8, and 5.5 mg/L
Measured concentrations as mg/L	Control: 0 hrs, 0.02; 48 hrs <0.02 mg/L 0.2 mg/L: 0 hrs, 0.022; 48 hrs 0.05 mg/L 5.50 mg/L: 0 hrs, 5.12; 48 hrs, 2.94 mg/L
Unit	mg/L
EC50, EL50, LC0, at 24,48 hours	48-hr EC0=0.4 mg/Land 48 hr EC50=1.1 mg/L; EC100=2.8 mg/L
Biological observations	The number of daphnids exhibiting immobilization at 48 hours for duplicate runs at each mean measured concentration was:0 mg/L, 0/40 & 0/40; 0.20 mg/L, 0/40 & 0/40; 0.4 mg/L, 0/40 0/40; 0.7, 4/40 & 5/40; 1.4 mg/L, 25/40 & 31/40; 2.8 mg/L, 40/40 & 32/40;
Control response satisfactory?	yes
Appropriate statistical evaluations?	None
Remarks fields for results	The respective ranges for pH, dissolved oxygen, and temperature were: 7.8-7.9, 8.1-8.5 mg/L, and 20.3-20.5C, respectively.
Conclusion remarks	The acute 48-hour EC50 for amylcinnamaldehyde in Daphnia magna under static conditions was 1.1 mg/L, respectively. The NOEC in Daphnid magna was 0.4 mg/L

Reliabilities	Reliability Code No. 1. Reliable without restriction.
Remarks for Data Reliability	The data are obtained by a recognized guideline method and are consistent with chemical structure.
References	Caspers (1992) Acute toxicity of amylcinnamaldehyde and the Daphnid, Daphnia magna. Study No. HR91/613101. Private communication to RIFM. Unpublished Report.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Remarks for Substance	Assay: 65-75%
Method/guideline	OECD 202-I Guideline (1984)
Test Type	Experimental (48- hr static test)
GLP	Yes
Year	1999
Analytical procedures	GLC
Species/Strain	Daphnia magna/Straus
Test details	Daphnia (2-24 hrs. old) were decanted into 25 ml glass beakers, each containing 10 ml of test solution with the test substance in various concentrations. Test solutions were prepared by emulsification of the test substance in water with acetone. There were 5 Daphnia per beaker and 5 beakers per each concentration. Test conditions consisted of a 16 hr./8 hr. light/dark cycle, a light intensity of 200 lx, oxygen concentration of 8.4-8.6 and temperature of 20.2-20.5 degrees Celsius. Test solutions were exposed to 16 hours of light and 8 hours of darkness. The Daphnia are examined for mobility after 24 and 48 hours. Daphnia which showed no reaction after 15 seconds were considered immobile. The pH, oxygen content and temperature were measured at the beginning and end of the test. Probit analysis was performed to determine the EC50.
Remarks for Test Conditions	None
Nominal concentrations as mg/L	0, 3.8, 7.5, 15.0, 30.0, and 60 mg/L
Measured concentrations as mg/L	
Unit	mg/L
EC50, EL50, LC0, at 24,48 hours	48 hr EC50=11.5 mg/L; EC100=30.0 mg/L
Biological observations	The number of daphnids exhibiting immobilization at 48 hours for quadruplicate runs at each mean measured concentration was:0, 3.8, 7.5 mg/L, 0/5 for all tests; 15 mg/L, 4/5 (3 runs) & 5/5 (1 test); 30 and 60 mg/L 5/5 at in all tests.

Control response satisfactory?	yes
Appropriate statistical evaluations?	Probit Analysis
Remarks fields for results	The respective ranges for pH, dissolved oxygen, and temperature were: 7.75-8.19, 8.2-8.4 mg/L, and 20.2-20.5C, respectively.
Conclusion remarks	The acute 48-hour EC50 for cinnamaldehyde (65-75%) in Daphnia magna under static conditions was 11.5 mg/L.
Reliabilities	Reliability Code No. 1. Reliable without restriction.
Remarks for Data Reliability	The data are obtained by a recognized guideline method and are consistent with chemical structure.
References	Barth M. and Winkler J. (2001) Acute toxicity of cinnamon bark oil in Daphnia magna. OECD Guideline 202-1. Study No. 001048503. Private communication to RIFM. Unpublished Report.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Remarks for Substance	Assay: 100%
Method/guideline	OECD 202
Test Type	Experimental (48- hr semi-static test with 24 hour renewal)
GLP	Yes
Year	2003
Analytical procedures	HPLC/UV detector
Species/Strain	Daphnia magna/Aquatic Biosystems, Inc.
Test details	48 hrs.
Remarks for Test Conditions	Juvenile daphnids (<24 hours old) produced from an in-house culture of adults were maintained at the contract laboratory under test conditions for 45 days. During the 48 hours prior to testing, the daphnid culture was maintained in 100% dilution water under static, renewal conditions for 48 hours. There was no mortality during the 48 hours prior to test and the test organisms appeared free of disease, injuries, or abnormalities. The daphnid culture produced young before day 12 and a subsample of adults produced on average, more than 3 young per day during the 7days prior to the beginning of the test. Ten daphnid were randomly selected for each replicate test. Tests were performed at 5 nominal concentrations. During the 48-hr test, daphnid were exposed to 16 hours of light and 8 hours of darkness. Mortality, immobility, and sub-lethal effects were

	determined visually at 0, 24, and 48 hours. Test solutions were renewed every 24 hours. Test temperature was maintained at 19.5-20.7 °C
Nominal concentrations as mg/L	0, 2.0. 3.3, 5.5, 9.0 15, and 25 mg/L
Measured concentrations as mg/L	0.00452 (control), 1.91, 3.34, 5.30, 9.57, 13.9, and 25.5 mg/L (mean concentration)
Unit	mg/L
EC50, EL50, LC0, at 24,48 hours	48-hr EC50=3.86 mg/Land 48 hr LC50=4.22 mg/L; NOEC 1.91 mg/L
Biological observations	The number of surviving daphnids at 48 hours for duplicate runs at each mean measured concentration was:0 mg/L, 10/10 & 10/10; 1.91 mg/L, 9/10 & 10/10; 3.34 mg/L, 9/10 8/10; 5.30 mg/L, 2/10 & 1/10; 9.57 mg/L, 1/10 & 0/10; 13.9 and 25.5 mg/L 0/10 & 0/10.
Control response satisfactory?	yes
Appropriate statistical evaluations?	Probit method (Stephan, 1978)
Remarks fields for results	The measured concentrations after 24 and 48 hours were 93-106% of the nominal concentrations, with the concentration being held steady throughout the test period. The respective ranges for conductivity, pH, dissolved oxygen, and temperature were: 590-600 umhos/cm, 6.8-7.1, 8.2-9.1 mg/L, and 19.5-20.7C, respectively.
Conclusion remarks	The acute 48-hour EC50 and LC50 for cinnamaldehyde in Daphnid magna under semi-static conditions were 3.86 and 4.22 mg/L, respectively. The NOEC for cinnamaldehyde in Daphnid magna is 1.91 mg/L
Reliabilities	Reliability Code No. 1. Reliable without restriction.
Remarks for Data Reliability	The data are obtained by a recognized guideline method and are consistent with chemical structure.
References	Ward T. (2003a) Acute toxicity test with cinnamaldehyde and the Daphnid, Daphnia magna. Study No. 2463-FF. Private communication to FFHPVC. Unpublished Report.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	ECOSAR
Test Type	Calculated based on log Kow
Species/Strain	Daphnia magna
Test Details	48 hrs
Remarks for Results	LC50 = 8.1  mg/l
Remarks for Data Reliability References	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.  ECOSAR

Substance Name	alpha-Amylcinnamaldehyde
CAS No.	122-40-7

Method/guideline ECOSAR

**Test Type** Calculated based on measured Kow

Species/Strain Daphnia magna

Test Details 48 hrs

Remarks for Results LC50 = 0.42 mg/l

and are consistent with chemical structure. Data are considered

reliable.

References ECOSAR

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0

Method/guideline ECOSAR

**Test Type** Calculated based on measured Kow

Species/Strain Daphnia magna

Test Details 48 hrs

Remarks for Results LC50 = 0.22 mg/l

and are consistent with chemical structure. Data are considered

reliable.

References ECOSAR

Substance Name	p-tert-Butyl-alpha-methyldihydrocinnamic aldehyde
CAS No.	80-54-6
Method/guideline	ECOSAR

**Test Type** Calculated based on measured Kow

Species/Strain Daphnia magna

Test Details 48 hrs

Remarks for Results LC50 = 0.40 mg/l

The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered Remarks for Data Reliability

reliable.

References **ECOSAR** 

#### 3.3 **Acute Toxicity to Aquatic Plants**

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Remarks for Substance	Assay: 100%
Method/guideline	OECD 201 Guideline
Test Type	Experimental
GLP	Yes
Year	2003
Species/Strain/Supplier	Green algae/Selenastrum capricornutum/UTEX 1648
Exposure period (duration)	72 hrs
Analytical monitoring	HPLC/UV detector
Remarks for Test Conditions  Nominal concentrations as	Green Algae/Selenastrum capricornutum/U. of Texas was maintained at test conditions for 14 days prior to the test. The culture was growing in at least 2 subcultures prior to the initiation of the test. In a range finding test, the number of cells/mL was >100 % of controls at 0.10 mg/L, 80% at 1.0 mg/L, and <1% at 100 mg/L after three days. In the definitive test, algae was treated with nominal concentrations of 0, 0.50, 1.0, 2.0, 4.0 and 8.0 mg/L for 72 hours. pH was adjusted to 7.5 and solutions were exposed for 24 hours of light of intensity, 400-410 foot candles. The number of algal cells/mL as well as relative size, cell shapes, color, adherence and aggregation of cells was determined. At 24, 48, and 72 hours 3 treatment and 6 control vessels were sacrificed to determine the number of algal cells/mL. Concentrations were determined by HPLC. 0, 0.5, 1.0, 2.0, 4.0, and 8.0 mg/L
mg/L	
Measured concentrations as mg/L	Initial mean measured concentrations 0.523, 1.04, 2.00, 3.80, and 7.03 mg/L; Final measured were 88-105% of nominal concentrations
Unit	mg/L
NOEC, LOEC or NOEL, LOEL	72 hr EC50=6.87 mg/L based on average specifc growth rate; 72-hr EC50=4.56 mg/L calculated using the number of cells/mL; 72-hr EC50= 4.07 mg/L using the area under the

	growth curve. The 72-hr NOEC=2.00 mg/L based on number of cells/mL			
Biological observations	Control algal populations grew at an acceptable rate (366,000 cells/ml) after 72 hours. Incubation temperatures were in the range from 23.2 to 24.0 C over the 72 hours and pH was unchanged by the test substance. At the conclusion of the test, samples of test media from each test vessel with maximal growth inhibition were combined with fresh media. After 48 hours incubation the number of cells increased from 850 cells/mL to 68,000 cells/mL at 7.03 mg/L suggesting that the toxic effects were algistatic.			
Appropriate statistical evaluations?	EC50 values determined by weighted least squares non-linear regression (Bruce and Versteeg, 1992); NOEC was determined using a one-way analysis of variance (ANOVA) and Bonferroni's test (Gulley et al. 1990)			
Conclusion remarks	The acute toxicity of cinnamaldehyde measured as a 50% decrease in growth and reproduction of freshwater algae was estimated to be 72 hr EC50=6.87 mg/L based on average specifc growth rate; 72-hr EC50=4.52 mg/L calculated using th number of cells/mL; 72-hr EC50= 4.07 mg/L using the area under the growth curve. The 72-hr NOEC=2.00 mg/L			
Reliabilities	Relabitiy code 1. Reliable without restrictions.			
Remarks for Data Reliability	OECD 201 Guideline study			
References	Ward T. (2003b) The growth and reproduction toxicity test with cinnamaldehyde and freshwater alga, Selenastrum capricornutum. OECD 201. Study No. 2464-FF. Private Communication to FFHPVC. Unpublished Report.			
Substance Name	alpha-Amylcinnamaldehyde			
CAS No.	122-40-7			
Remarks for Substance	Assay:			
Method/guideline	OECD 201 Guideline			
Test Type	Experimental			
GLP	Yes			
Year	2003			
Species/Strain/Supplier	Green algae/Selenastrum capricornutum/UTEX 1648			
Exposure period (duration)	72 hrs			
Analytical monitoring	HPLC/UV detector			
Remarks for Test Conditions	Green Algae/Selenastrum capricornutum/U. of Texas was maintained at test conditions for 14 days prior to the test. The culture was growing in at least 2 subcultures prior to the initiation of the test. In a range finding test, the number of cells/mL was 83 % of controls at 0.10, 46% at 1.0 mg/L, and			

	<1% at 10 mg/L and 100 mg/L after three days. In the definitive test, algae was treated with nominal concentrations of 0, 0.095, 019, 0.38, 0.75, 1.50 and 3.0 mg/L for 72 hours. pH was adjusted to 7.5 and solutions were exposed for 24 hours of light of intensity, 400-410 foot candles. The number of algal cells/mL as well as relative size, cell shapes, color, adherence and aggregation of cells was determined. At 24, 48, and 72 hours 3 treatment and 6 control vessels were sacrificed to determine the number of algal cells/mL. Concentrations were determined by HPLC.	
Nominal concentrations as mg/L	0, 0.095, 019, 0.38, 0.75, 1.50 and 3.0 mg/L	
Measured concentrations as mg/L	Initial mean measured concentrations 0.0934, 0.154, 0.363, 0.651, 1.39, and 2.75 mg/L; Final measured were 81-98% of nominal concentrations	
Unit	mg/L	
NOEC, LOEC or NOEL, LOEL	72 hr EC50=1.88 mg/L based on average specifc growth rate; 72-hr EC50=1.18 mg/L calculated using the number of cells/mL; 72-hr EC50= 1.24 mg/L using the area under the growth curve. The 72-hr NOEC=0.154 mg/mL based on number of cells/mL, average growth rate, or area under the growth curve.	
Biological observations	Control algal populations grew at an acceptable rate (312,000 cells/ml) after 72 hours. Incubation temperatures were in the range from 23.7 to 24.0 C over the 72 hours and pH was unchanged by the test substance. At the conclusion of the test, samples of test media from each test vessel with maximal growth inhibition were combined with fresh media. After 48 hours incubation the number of cells increased from 300 cells/mL to 140,000 cells/mL at 2.95 mg/L suggesting that the toxic effects were algistatic.	
Appropriate statistical evaluations?	EC50 values determined by weighted least squares non-linear regression (Bruce and Versteeg, 1992); NOEC was determined using a one-way analysis of variance (ANOVA) and Bonferroni's test (Gulley et al. 1990)	
Conclusion remarks	The acute toxicity of methylionone measured as a 50% decrease in growth and reproduction of freshwater algae was estimated to be 72 hr EC50=1.88 mg/L based on average specifc growth rate; 72-hr EC50=1.18 mg/L calculated using the number of cells/mL; 72-hr EC50= 1.24 mg/L using the area under the growth curve. The 72-hr NOEC=0.154 mg/L	
Reliabilities	Relabitiy code 1. Reliable without restrictions.	
Remarks for Data Reliability	OECD 201 Guideline study	
References	Ward T. (2003c) The growth and reproduction toxicity test with alpha-amylcinnamaldehyde and freshwater alga, Selenastrum capricornutum. OECD 201. Study No. 2462-FF. Private Communication to FFHPVC. Unpublished Report.	

Substance Name	Cinnamaldehyde	

**CAS No.** 104-55-2

Method/guideline Test compounds were dissolved under sterile conditions in

modified KNOP solution. Subsequently, these solutions were measured into a flask to which a growing cell suspension was added. Cultures were shaken for 48 hr, where after the cells

were centrifuges.

Species/Strain/Supplier Chlorella vulgaris

**Exposure period** 96 hrs

Remarks for Test Conditions After acidification to 4.0 agueous solution was extracted w/

ether. Ether fractions were treated w/anhydrous sodium sulfate, filtered & concentrated. Ethanol was added to obtain a final extract of 1 ml. From this extract, a sample was subjected to

TLC.

**Biological Observations**Cinnamic aldehyde was found to inhibit the algae growth in a

concentration as low as  $5X10(-5)\,M$ . At the same concentration a stimulation of the respiration of the algae was observed at pH

5.6 & pH 7.2

**Conclusion Remarks**Cinnamic aldehyde inhibited the algal growth and stimulated

the respiration.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Dedoner, A. and VanSumere, C.F. (1971). The effect pf

phenolics and related compounds on the growth and the respiration of Chlorella vulgaris. Z. PflPhysiol 65(1): 70-80.

Substance Name alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Method/guideline Calculated

Test Type ECOSAR

Species/Strain/Supplier Green algae

**Exposure period** 96 hrs

**Conclusion Remarks** EC50 = 0.87 mg/l

Remarks for Data Reliability The data are obtained by a recognized SAR calculation method

and are consistent with chemical structure. Data are considered

reliable.

References ECOSAR

Substance Name	alpha-Hexylcinnamaldehyde		
CAS No.	101-86-0		
Method/guideline	Calculated		
Test Type	ECOSAR		

Species/Strain/Supplier Green algae

**Exposure period** 96 hrs

**Conclusion Remarks** EC50 = 0.34 mg/l

and are consistent with chemical structure. Data are considered

reliable.

References ECOSAR

Substance Name	p-tert-Butyl-alpha-methyldihydrocinnamic aldehyde		
CAS No.	80-54-6		
Method/guideline	Calculated		
Test Type	ECOSAR		
Species/Strain/Supplier	Green algae		
Exposure period	96 hrs		
Conclusion Remarks	EC50 = 0.827  mg/l		
Remarks for Data Reliability	The data are obtained by a recognized SAR calculation method and are consistent with chemical structure. Data are considered reliable.		
References	ECOSAR		

## 4 Human Health Data

## 4.1 Acute Toxicity

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	
Method/guideline	LD50 was computed by method of Litchfield & Wilcoxon (1949).	
Test Type	Acute Oral LD50	
GLP	Not reported	
Year	1964	
Species/Strain	Guinea pig	
Sex	Male and Female	

# of animals per sex per

dose

NG

Route of administration O

Oral

Value LD50 or LC50 with

confidence limits Remarks for Results LD50 = 1160 (95% CI 950-1420) mg/kg.

The LD50 was reported to be 1160 mg/kg. Coma was reported

with higher doses.

**Conclusion Remarks** The LD50 was reported to be 1160 (95%Cl 950-1420) mg/kg.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal Food

Cosmetic Toxicology,

**References** Jenner, P. M., Hagan, E.C., Taylor, J.M, Cook, E.L. and

Fitzhugh, O.G. (1964). Food Flavorings and Compounds of Related Structure I. Acute Oral Toxicity. Food and Cosmetics

Toxicology 2(3): 327-343.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	

Method/guideline A group of animals, 6 animals per group per sex were given the

test substance by oral gavage.

**Test Type** Acute Oral LD50 test

**GLP** NG

**Year** 1974

Species/Strain Rat/White

**Sex** Male and Female

# of animals per sex per

dose

6

Vehicle Sunflower oil

Route of administration Oral (gavage)

Remarks for Test Conditions No other details were given

Value LD50 or LC50 with confidence limits

LD50 = 3400 mg/kg or 25.8 mM.

Remarks for Results No other details were given

**Conclusion Remarks** The oral LD50 value for cinnamaldehyde was calculated to be

3400 mg/kg

**Data Qualities Reliabilities** Reliability code 3. Data not reliable. The data must be viewed

with caution.

Remarks for Data Reliability Original article is in Russian. English translation doesn't report

details or these details are missing in the original article.

General Remarks Authors claim that the acute oral LD50 values for

Cinnamaldehyde for rats, mice and guinea pigs was the same

value of 3400 mg/kg.

Zaitsev, A, N. and Rakhmanina (1974). Some Data on the Toxic Properties of Phenylethyl and Cinnamyl Alcohol Derivatives. Vopr. Pitaniya 6: 48-53. References

Substance Name	Cinnamaldehyde
Substance Name	Cililamateriyde
CAS No.	104-55-2
Method/guideline	A group of animals, 6 animals per group per sex were given the
Test Type	test substance by oral gavage. Acute Oral LD50 test
GLP	NG
Year	1974
Species/Strain	Mice/White
Sex	Male and Female
# of animals per sex per	6
Vehicle	Sunflower oil
Route of administration	Oral (gavage)
Remarks for Test Conditions	No additional details given.
Value LD50 or LC50 with confidence limits	LD50 = 3400 mg/kg or 25.8 mM.
Remarks for Results	No other details were given
Conclusion Remarks	The oral LD50 value for cinnamaldehyde was calculated to be
Data Qualities Reliabilities	3400 mg/kg Reliability code 3. Data not reliable.
Remarks for Data Reliability	Original article was in Russian. English translation either doesn't report details or these details are missing in the original article.
General Remarks	Authors claim that the acute oral LD50 values for Cinnamaldehyde for rats, mice and guinea pigs was same value of 3400 mg/kg.
References	Zaitsev, A, N. and Rakhmanina (1974). Some Data on the Toxic Properties of Phenylethyl and Cinnamyl Alcohol Derivatives. Vopr. Pitaniya 6: 48-53.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	
Method/guideline	A group of animals, 6 animals per group per sex were given the test substance by oral gavage.	
Test Type	Acute Oral LD50 test	
GLP	NG	

**Year** 1974

Species/Strain Guinea pig

**Sex** Male and Female

# of animals per sex per

dose

6

Vehicle Sunflower oil

Route of administration Oral (gavage)

Remarks for Test Conditions No additional details given.

Value LD50 or LC50 with

confidence limits Remarks for Results LD50 = 3400 mg/kg or 25.8 mM.

No other details given

**Conclusion Remarks** The oral LD50 value for cinnamaldehyde was calculated to be

3400 mg/kg

**Data Qualities Reliabilities** Reliability code 3. Data not reliable. The data must be viewed

with caution.

Remarks for Data Reliability Original article was in Russian. English translation either

doesn't report details or these details are missing in the original

article

General Remarks Authors claim that the acute oral LD50 values for

Cinnamaldehyde for rats, mice and guinea pigs was same

value of 3400 mg/kg.

**References** Zaitsev, A, N. and Rakhmanina (1974). Some Data on the

Toxic Properties of Phenylethyl and Cinnamyl Alcohol

Derivatives. Vopr. Pitaniya 6: 48-53.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	

Method/guideline The study was performed on albino rabbits according to the

method described under section 191.10 of the final order enforcement Regulation, Federal Register Vol. 26, No. 155,

p7336, Aug 12, 1961.

**Test Type** Acute Dermal LD50

GLP Not reported

**Year** 1973

Species/Strain Rabbit/Albino

Sex Not reported

# of animals per sex per

dose

20

Vehicle None reported

Route of administration Dermal

**Remarks for Test Conditions** The test substance was applied to the intact or abraded skin of

> the rabbit. The mortality data was evaluated according to the Thompson moving method as described by Carrol S. Weil. Biometrics 8(3): 249-263, 1952. Doses tested 0.25, 0.50, 1.0,

2.0 & 4.0 ml/kg.

Value LD50 or LC50 with

confidence limits

**Conclusion Remarks** 

Number of deaths at each

**Data Qualities Reliabilities** 

dose level

Acute Dermal LD50 & 19/20 Confidence Limit = 0.59 (0.42-

0.84) ml/kg. LD50=620 mg/kg bw.

0.25 ml/kg- 0/2 death (Intact or abraded); 0.50 ml/kg- 1/2 deaths in abraded group; 1.0 ml/kg- 2/2 deaths in both intact & abraded group; 2.0 ml/kg- 2/2 deaths in both intact and abraded group; 4.0 ml/kg- 2/2 deaths in both intact & abraded group. Cinnamic aldehyde has an acute dermal LD50 and 19/20

Confidence limits of 0.59 0.42-0.884) ml/kg.

Reliability code 1. Reliable without restrictions.

Shelanski, M. and Moldovan, M. (1973). Report to RIFM by References

Food and Drug Research Laboratories. Feb 16, 1973.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	
Method/guideline Test Type	Rabbits were dosed dermally with cinnamic aldehyde at 0.59, 0.83, 1.00, 1.23 & 1.50 ml/kg. The test substance was kept in contact with the skin for 24 hours. The animals were observed daily for signs of mortality, toxicity and pharmacological effects. Acute Dermal Toxicity	
GLP	Yes	
Year	1986	
Species/Strain	New Zealand Albino rabbits	
Sex	Male and Female	
# of animals per sex per	4	
dose Route of administration	Dermal	
Remarks for Test Conditions	Skin reactions were scored on days 1, 7 and 14. Body weights were recorded pretest and at termination. All animals were examined for gross pathology. The LD50 was calculated by the method of Litchfield and Wilcoxon.	

Value LD50 or LC50 with confidence limits

Number of deaths at each dose level

The LD50 and 95% confidence limits are: 1.2 (0.9 - 1.6) ml/kg of the body weight.

0.59 ml/kg= 0 dead/2 treated; 0.83 ml/kg = 2 dead/4 treated; 1.00 ml/kg = 1 dead/4 treated; 1.23 ml/kg = 1 dead/4 treated;

1.50 ml/kg = 4 dead/4 treated.

**Remarks for Results** Deaths occurred by day 3, and were preceded with predeath

physical signs of few feces, lethargy, ataxia and rales. Necropsy of the deaths revealed abnormalities of the lungs, liver, kidneys, treated skin and GI tract, as well as brown staining of the anogenital area and yellow staining of the nose/mouth area. Survivors: signs of diarrhea, few feces, emaciation, ataxia and limited mobility due to severe skin

reaction, a	abnormalities	of skin a	and intestines.	Larger than
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normal uterus.

**Conclusion Remarks** The LD50 and 95% confidence limits are: 1.2 (0.9 - 1.6) ml/kg

of the body weight. LD50=1260 mg/kg bw

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was conducted in accordance with GLP

Fritzsche Dodge and Olcott, Inc. (1986). Acute Dermal Toxicity References

of Cinnamaldehyde in Rabbits. Unpublished. Report to RIFM.

Substance Name	p-t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	Acute oral toxicity was determined in rats.
Test Type	Acute Oral Toxicity
GLP	NG
Year	1977
Species/Strain	Rats

Not reported Sex

# of animals per sex per

dose

10

Oral Route of administration

**Remarks for Test Conditions** Doses used: 1.22, 2.47, 5.0 and 10.14 g/kg

g/kg

10/10

Value LD50 or LC50 with

confidence limits

Number of deaths at each dose level

Remarks for Results

**Conclusion Remarks** 

Toxic signs = 1.22 g/kg: diarrhea; 2.47 g/kg: piloerection,

lethargy, flaccid; 5.0 g/kg: lethargy, piloerection, diarrhea. coma; 10.14 g/kg: ataxia, lethargy, piloerection and diarrhea. The oral LD50 and 95% confidence interval are 3.7 (2.6-5.4)

The oral LD50 and 95% confidence interval are 3.7 (2.6-5.4)

1.22 g/kg = 0/10; 2.47 g/kg = 1/10; 5.0 g/kg = 7/10; 10.14 g/kg = 1/10

g/kg. LD50=3700 mg/kg bw.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

References Moreno O. M. (1977b). Acute Oral toxicity in Rats. Dermal

Toxicity in Rabbits. Unpublished. Report to RIFM.

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	40 Male Wistar rat strain were used. Rats were observed for signs of toxicity and pharmacologic effect at 1, 6 & 24 hours and daily thereafter for a period of 14 days.
Test Type	Oral LD50

**GLP** NG

Year 1971

Species/Strain Male Wistar rats

Sex Male

# of animals per sex per

10

dose

Route of administration Oral

**Remarks for Test Conditions** Doses tested: 1.78, 2.67, 4.0 and 6.0 gm/kg.

Value LD50 or LC50 with

confidence limits

LD50 (95% Confidence Limit) = 3.1 (3.75-2.45) g/kg

Number of deaths at each

dose level

1.78 g/kg = 1/10; 2.67 g/kg = 4/10; 4.0 g/kg = 7/10; 6.0 g/kg =

10/10.

**Remarks for Results** Symptomology: Depression, Lethargy, Anorexia, Weight loss

**Conclusion Remarks** The oral LD50 was reported to be 3.1 g/kg. LD50=3100 mg/kg

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

References Moreno O.M. (1971) Acute toxicity studies in rats, mice, rabbits

and guinea pigs. Unpublished report to RIFM.

**Substance Name** Cinnamaldehyde

CAS No. 104-55-2

Method/guideline Acute Dermal toxicity

**Test Type** Acute Dermal LD50

**GLP** Not reported

Year 1972

Species/Strain Rabbits

Sex Not reported

# of animals per sex per

dose

6

Vehicle Not reported

Route of administration Dermal

**Remarks for Test Conditions** Dose tested = 5.0 g/kg

Value LD50 or LC50 with

confidence limits

Dermal LD50 <5.0 g/kg.

Number of deaths at each

All animals died overnight after dosing.

dose level

**Conclusion Remarks** The dermal LD50 value for cinnamic aldehyde in rat is less than

5 g/kg.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

References

Shelanski M. and Moldovan, M. (1973). Report to RIFM by Food and Drug Research Laboratories. Feb 16, 1973. Shelanski, M. and Moldovan, M. (1973). Report to RIFM by Food and Drug Research Laboratories. Feb 2, 1973.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	
Method/guideline Test Type	Rabbits were dosed dermally with 1000 mg/kg of the test material and kept in contact with the skin for 24 hours. Derma responses were recorded 24 hours, day 7 and 14 postdose. Dermal LD50	
• •		
GLP	Yes	
Year	1996	
Species/Strain	New Zealand White rabbits	
Sex	Male and Female	
# of animals per sex per dose	5	
Vehicle	NG	
Route of administration	Dermal	
Remarks for Test Conditions	Body weights were recorded pretest and at death, or termination in the survivors. All animals were examined for gross pathology. The test sites were scored using the numerical Draize scoring code. An estimate of the LD50 was made based on the survival during the study.	
Value LD50 or LC50 with confidence limits	The LD50 is greater than 1000 mg/kg of body weight.	
Number of deaths at each dose level	All animals survived the 1000 mg/kg dermal application.	
Remarks for Results	Necropsy revealed treated skin abnormalities in all animals. Liver abnormalities were noted in one animal, and kidney abnormalities in three animals, one of which had wetness of the anogenital area.	
<b>Conclusion Remarks</b>	The dermal LD50 was reported to be greater than 1000 mg/kg.	
Data Qualities Reliabilities	Reliability code 1. Reliable without restrictions.	
Remarks for Data Reliability	The study was conducted in accordance with GLP.	
References	MB Research Labs (1996) Unpublished Report to RIFM.	

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	

Method/guideline Oral LD50

**Test Type** Acute Oral LD50

GLP Not reported

**Year** 1965

Species/Strain Rats

Sex Not reported

# of animals per sex per

dose

NG

Vehicle NG

Route of administration Oral

**Remarks for Test Conditions** Article in Romanian. Details not given in the English abstract.

Value LD50 or LC50 with

confidence limits

LD50 = 3350 mg/kg.

Number of deaths at each

dose level

Article in Romanian. Details not given in the English abstract.

Remarks for Results Article in Romanian. Details not given in the English abstract.

Conclusion Remarks LD50 = 3.350 mg/kg

**Data Qualities Reliabilities** Reliability code 3. Data not reliable.

Remarks for Data Reliability Article in Romanian. Details not given in the English abstract.

**References** Sporn A. (1965). Investigation of the Toxicity of Cynamic

Aldehyde. Igiena 14(6): 339-346.

Substance Name	Cinnamaldehyde

**CAS No.** 104-55-2

Method/guideline LD50

**Test Type** Intraperitoneal LD50

GLP Not reported

**Year** 1965

Species/Strain Mice

Sex Not reported

# of animals per sex per

dose

NG

Vehicle NG

Route of administration Intraperitoneal

Remarks for Test Conditions Article in Romanian. Details not given in the English abstract.

Value LD50 or LC50 with confidence limits

LD50 = 2318 mg/kg.

confidence limits

Remarks for Results

Article in Romanian. Details not given in the English abstract.

**Conclusion Remarks** Intraperitoneal LD50 for Cinnamaldehyde in mice was shown to

be 2318 mg/kg.

**Data Qualities Reliabilities** Reliability code 3. Data not reliable.

Remarks for Data Reliability Article in Romanian. Details not given in the English abstract.

**References** Sporn A. (1965). Investigation of the Toxicity of Cynamic

Aldehyde. Igiena 14(6): 339-346.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	LD50 was computed by method of Litchfield & Wilcoxon (1949).
Test Type	Acute Oral LD50
GLP	Not reported
Year	1964
Species/Strain	Osborne-Mendel rats
Sex	Male and Female
# of animals per sex per	10
dose Vehicle	None
Route of administration	Oral

Value LD50 or LC50 with	LD50 = 2220 (1910-2600) mg/kg
confidence limits	

NG

Number of deaths at each

dose level

Remarks for Results The LD50 was reported to be 2220 mg/kg. Depression,

diarrhea and scrawny appearance were noted.

**Conclusion Remarks** The LD50 was reported to be 2220 (1910-2600) mg/kg.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal Food

Cosmetic Toxicology,

References Jenner, P. M., Hagan, E.C., Taylor, J.M, Cook, E.L. and

Fitzhugh, O.G. (1964). Food Flavorings and Compounds of Related Structure I. Acute Oral Toxicity. Food and Cosmetics

Toxicology 2(3): 327-343.

Substance Name	alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Method/guideline** Six rabbits were used at 3 different dose levels 1, 2 & 3 g/kg.

Chemical was applied to clipped area and was occluded for 24

hrs and the animals were observed for 7 days.

**Test Type** Acute Dermal Toxicity test

**GLP** NG

**Year** 1971

Species/Strain Rabbits

Sex Female

# of animals per sex per

dose

2

Route of administration Dermal

Remarks for Test Conditions Highest dose was limited by the area available for treatment as

well as by the chemical available.

Value LD50 or LC50 with

confidence limits

Number of deaths at each

dose level

greater than 3 g/kg. No animals died at any dose level tested.

Remarks for Results Moderate erythema was seen. Occasional sloughing was seen

but this was in large part due to damage caused by the removal

The dermal lethal dose of the test substance was reported to be

of the tape from the skin.

Conclusion Remarks The dermal lethal dose of the test substance was reported to

greater than 3 g/kg. Dermal LD50>3000 mg/kg bw.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**References** Moreno O.M. (1971) Acute toxicity studies in rats, mice, rabbits

and guinea pigs. Unpublished report to RIFM.

**Substance Name** p-t-Butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Method/guideline** Acute dermal toxicity was determined in rabbits.

**Test Type** Acute Dermal Toxicity

GLP NG

**Year** 1977

Species/Strain Rabbits

Sex Not reported

# of animals per sex per

dose

10

Vehicle Not reported

Route of administration Dermal

Value LD50 or LC50 with

confidence limits

Number of deaths at each

dose level

than 5 g/kg.

No death occurred

Remarks for Results

Mild redness was seen in 4 animals; moderate redness in 6 animals, mild edema in 7 animals and moderate edema in 3

The dermal LD50 value for Lilial was calculated to be greater

animals.

**Data Qualities Reliabilities** 

Reliability code 1. Reliable without restrictions.

Conclusion Remarks

The dermal LD50 value for Lilial was calculated to be greater

than 5000 mg/kg.

References

Moreno O. M. (1977b). Acute Oral toxicity in Rats. Dermal

Toxicity in Rabbits. Unpublished. Report to RIFM.

alpha-Hexylcinnamaldehyde, light brown liquid with aromatic **Substance Name** odor

CAS No. 101-86-0

Method/quideline 5 Sprague-Dawley rats per sex per dose received a single 4 hr

exposure to aerosol containing test substance. Animals were observed for 14 days for body weight changes, mortality, clinical signs, gross and histopathological changes.

Acute Inhalation toxicity **Test Type** 

**GLP** NG

1980 Year

Species/Strain Sprague-Dawley rats

Male and Female Sex

# of animals per sex per

dose

5

Route of administration Inhalation

**Remarks for Test Conditions** The nominal chamber concentration, calculated from airflow

> and quantity of test article consumed was 5.00 mg/L. The mean value for the measured concentration was 2.12 mg/L in the

chamber.

LC50> 5 mg/L

Value LD50 or LC50 with

confidence limits Number of deaths at each

dose level

No deaths were reported

Remarks for Results Enlarged bronical lymph nodes sometimes accompanied by

> pulmonary congestion, multiple grey-green pinpoint foci in the lungs, minimal loss of body weight on the days immediately

following treatment.

Conclusion Remarks The acute median lethal concentration was calculated to be

greater than 5.00 mg/L expressed in terms of nominal

concentration.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

References Breckenridge C. (1980). The acute toxicity of inhaled hexyl

cinnamic aldehyde in the albino rats. Unpublished. Report to

RIFM.

Out stance No.	A D. C. Later and A. L. Land	
Substance Name	<i>p-</i> t-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde, clear liquid.	
CAS No.	80-54-6	
Method/guideline  Test Type	Test substance was applied at a dose of 5 ml/kg to the shaved skin of three rabbits of each sex and occluded for 24 h after which the rabbits were observed for 14 days for overt toxic signs and mortality.  Acute Dermal Toxicity	
• •	·	
GLP	Yes	
Year	1979	
Species/Strain	Albino New Zealand rabbits	
Sex	Male and Female	
# of animals per sex per dose	3	
Route of administration	Dermal	
Remarks for Test Conditions		
Value LD50 or LC50 with confidence limits Number of deaths at each dose level Remarks for Results	Dermal LD50 > 5 ml/kg  There were no deaths.  Treatment caused moderate erythema and thickened, wrinkled skin in all test animals, persisting through Day 9. Subcapsular	
Conclusion Remarks  Data Qualities Reliabilities	(agonal) hemorrhages of the kidneys were found at necropsy in most of the test animals.  The acute dermal LD50 for the test substance was reported to be greater than 5 ml/kg. Acute dermal LD50>5250 mg/kg bw. Reliability code 1. Reliable without restrictions.	
Remarks for Data Reliability	The study was conducted in accordance with GLP.	
References	Slepetys (1979). Cosmopolitan Safety Evaluation Unpublished Report. FEMA 15027.	
Substance Name	alpha-Amylcinnamaldehyde	
CAS No.	122-40-7	
Method/guideline	Acute dermal toxicity was determined in rabbits.	
Test Type	Acute Dermal Toxicity	

NG

1973

Rabbits

GLP

Year

Species/Strain

Sex Not reported

# of animals per sex per

dose Vehicle 4

/ehicle Not reported

Route of administration Dermal

Value LD50 or LC50 with

confidence limits

Number of deaths at each

dose level

The dermal LD50 value for alpha-amylcinnamaldehyde was

calculated to be greater than 2000 mg/kg.

No death occurred

Remarks for Results The was no evidence of toxicity at 2000 mg/kg bw

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Conclusion Remarks The dermal LD50 value for alpha-amylcinnamaldehyde was

calculated to be greater than 2000 mg/kg.

**References** Moreno O. M. (1973). Acute Oral toxicity in Rats. Dermal

Toxicity in Rabbits. Unpublished. Report to RIFM.

Substance Name	alpha-Amylcinnamaldehyde
	,

**CAS No.** 122-40-7

Method/guideline LD50 was computed by method of Litchfield & Wilcoxon (1949).

**Test Type** Acute Oral LD50

GLP Not reported

**Year** 1964

Species/Strain Rst/Osborne-Mendel

**Sex** Male and Female

# of animals per sex per

dose

5

NG

Vehicle None

Route of administration Oral

Value LD50 or LC50 with confidence limits

Number of deaths at each

dose level

LD50 = 3730 ( 95% CI, 3190-4370 ) mg/kg. Slope function with

95% confidence interval=1.4 (1.2-1.6)

ose ievei

**Remarks for Results** 

The LD50 was reported to be 3730 mg/kg. Depression, porphyrin-like deposit around eyes and nose.

Conclusion Remarks The LD50 was reported to be 3730 (3190-4370) mg/kg.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal Food

Cosmetic Toxicology,

**References** Jenner, P. M., Hagan, E.C., Taylor, J.M, Cook, E.L. and

Fitzhugh, O.G. (1964). Food Flavorings and Compounds of

## 4.2 In Vitro Genotoxicity

Substance Name

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	
Method/guideline	Cinnamaldehyde was tested for its antimutagenic effect on mitomycin C pretreated cells.	
Test Type	Sister Chromatid Exchange	
System of Testing	Chinese Hamster ovary cell	
GLP	NG	
Year	1987	
Species/Strain	Chinese Hamster Ovary cells	
Doses/Concentration	0-20 uM	
Statistical Methods	NG	
Remarks for Test Conditions Results	Chinese hamster ovary cells were treated in fresh medium containing the mutagen for 22 h. After treatment, cells were washed & incubated with cinnamaldehyde for 22 h. BudR at 20 uM was added. Mitotic cells were collected by the addition of colchicine.  No increase in the frequencies of Sister Chromatid Exchange was observed after cells were treated with cinnamaldehyde alone. However, pretreatment of cells with mitomycin C resulted in increase in the frequency.	
Cytotoxic concentration	NG	
Genotoxic effects	None	
Statistical results	NG	
<b>Conclusion Remarks</b>	No evidence of mutagenicity by itself but increased the mutagenicity of mitomycin C.	
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.	
Remarks for Data Reliability	Study was published in a peer review journal.	
References	Sasaki, Y.F., Imanishi, H., Ohta, T. and Shirasu, Y. (1987), Effects of antimutagenic flavourings on SCEs induced by chemical mutagens in cultured Chinese hamster cells. Mutation Research 189: 313-318.	

Cinnamaldehyde

**CAS No.** 104-55-2

Method/guideline Ames

**Test Type** Reverse Mutation

System of Testing Bacterial

GLP NG

**Year** 1980

**Species/Strain** Salmonella typhimurium/TA100

Metabolic Activation With and without rat liver microsome fraction S9 from Aroclor

induced rats

**Doses/Concentration** 0.1, 0.2, 0.3 0.5, 1.2, 3 & 5 umoles/plate (13.2 to 320 ug/plate)

Statistical Methods NG

Results No mutagenic effects. Cinnamaldehyde and alpha-

methylcinnamaldehyde were non-mutagenic for Salmonella typhimurium TA100 both in the presence or absence of aroclor

1254 induced rat liver S9 mix.

Cytotoxic concentration NG

Genotoxic effects None

Statistical results NG

Remarks for Results Chloro or bromo substitution in the alpha-carbon position of

cinnamaldehyde leads to the derivatives that are strongly

mutagenic in Salmonella Typhimurium TA100.

Conclusion Remarks No mutagenic activity

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

**References** Neudecker, T., Ohrlein, K, Eder, E and Henschler, D. (1983).

Effect of Methyl and Halogen Substitutions in the *alpha* C position on the Mutagenicity of Cinnamaldehyde. Mutation

Research 110: 1-8.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2

**Method/guideline** Salmonella typhimurium strains TA97a, TA100, TA102 &

TA104 in the presence and absence of aroclor-induced liver

S9s from F344 rats & B6C3F1 mice.

**Test Type** Reverse Mutation

System of Testing Bacterial

**GLP** No

**Year** 1998

Species/Strain Salmonella typhimurium/TA97a, TA100, TA102 and TA104

Metabolic Activation With and without mice liver microsome fraction S9 from Aroclo

induced rats and mice.

**Doses/Concentration** 25, 50, 100, 200 and 300 ug/plate

Statistical Methods Dunnett's t-test and Wahrendorf ranking and linear regression.

**Remarks for Test Conditions** Positive control: 2-aminoanthracene.

**Results** trans-Cinnamaldehyde exhibited a weak mutagenic response in

TA100 with mouse liver S9 mix.

Cytotoxic concentration NG

**Genotoxic effects** Weak mutagenic response

Statistical results NG

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

**References** Dillon, D., Combes, R. and Zeiger, E. (1998). The Effectiveness

of Salmonella Strains TA100, TA102 and TA104 for Detecting Mutagenicity of Some Aldehydes and Peroxides. Mutagenesis

13(1): 19-26.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	

**Method/guideline** The mutagenicity test was conducted in the

Salmonella/microsome mutagenicity assay on plates according to the method of Ames with the Salmonella typhimurium TA98

and TA100.

**Test Type** Reverse Mutation

System of Testing Bacterial

**GLP** NG

**Year** 1982

Species/Strain Salmonella typhimurium TA 98, TA 100

Metabolic Activation Rat-liver microsome (S9) was prepared from Sprague-Dawley

rats treated with Aroclor 1254

**Doses/Concentration** 0.05 to 500 ug/plate.

Statistical Methods NG

Remarks for Test Conditions Diluted in DMSO

**Results** Negative

Cytotoxic concentration Not reported

**Genotoxic effects** Negative

Statistical results NG

**Conclusion Remarks** No mutagenic effects

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability Study was published in a peer review journal.

References Kasamaki, A., Takahashi, H., Tsumura, N., Niwa, J., Fujita, T.,

and Urasawa, S. (1982). Genotoxcity of Flavoring Agents.

Mutation Research 105:387-392

	Mutation Research 105:387-392.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline Test Type	Exponentially growing Chinese-hamster B241 were exposed to the test substance for 24 hr and then incubated for another 24 hrs without the test chemical followed by treatment with colchicine.  Chromosomal Aberration Test
System of Testing	Non Bacterial
GLP	NG
Year	1982
Species/Strain	Chinese hamster cell line B241
Metabolic Activation	Rat-liver microsome (S9) was prepared from Sprague-Dawley rats treated with Aroclor 1254. Rat-liver microsome (S9) was prepared from Sprague-Dawley rats treated with Aroclor 1254.
Doses/Concentration	Several doses up to 10 nM.
Statistical Methods	Chi-Square test
Remarks for Test Conditions	The test chemical was dissolved in DMSO at a concentration of

50 mM and then was diluted with the medium. Control cells were treated with a medium containing DMSO equal to the test

solution.

Results trans-Cinnamaldehyde exhibited high potential for inducing

> aberrations. The total frequency of the aberrations indicated a dose-dependent increase at a certain dose range. DMSO did not affect the frequency or the type of spontaneous aberrations

Cytotoxic concentration Not reported

Chromosomal aberrations Genotoxic effects

NG Statistical results

Remarks for Results Chromatid break, chromosome break, chromatid exchange,

ring or dicentric chromosomes, fragmentation, translocation and

pulverization were observed.

**Conclusion Remarks** Severe chromosome aberrations were observed in the cells

treated with Cinnamaldehyde,

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

**References** Kasamaki, A., Takahashi, H., Tsumura, N., Niwa, J., Fujita, T.,

and Urasawa, S. (1982). Genotoxcity of Flavoring Agents.

Mutation Research 105:387-392

	Mutation Research 105:387-392.
Substance Name	Cinnomoldobydo
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	The mutagenicity assay with Salmonella typhimurium was conducted as described by Ames et al with tester strain TA100 and TA98. S9 was prepared from the PCB-treated male Sprague-Dawley rats.
Test Type	Reverse Mutation assay
System of Testing	Bacterial
GLP	NG
Year	1982
Species/Strain	Salmonella typhimurium/TA100, TA98, TA1535, TA1537 and TA1538.
Metabolic Activation	S9 fraction was prepared from the PCB-treated male Sprague- Dawley rats
Doses/Concentration	60, 120, 300 and 600 ug/plate.
Statistical Methods	None performed

Remarks for Test Conditions Histidine-independent colonies were scored after incubation at

37C for 48-72 h.

Results No significant increase in revertant number with Salmonella

strains in the presence or absence of S9 fraction.

**Cytotoxic concentration** 600 ug/plate

Genotoxic effects Negative

Statistical results NG

Conclusion Remarks Cinnamaldehyde was not found to be mutagenic under the test

conditions

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Sekizawa, J. and Shibamoto, T. (1982). Genotoxicity of Safrole-

Related Chemicals in Microbial Test Systems. Mutation

Research 101: 127-140.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	The mutagenicity assay with E. coli WP2uorA trp- was performed according to the method described by Green and Murial (1976).
Test Type	Reverse Mutation
System of Testing	Bacterial
GLP	NG
Year	1982
Species/Strain	E. coli/WP2 uorA trp-
Metabolic Activation	S9 fraction was prepared from the PCB-treated male Sprague- Dawley rats
Doses/Concentration	60, 120, 300 and 600 ug/plate.
Statistical Methods	NG
Remarks for Test Conditions	After 48-72 h incubation at 37 °C, revertant colonies were counted.
Results	No mutagenic effects.
Cytotoxic concentration	600 ug/plate
Genotoxic effects	None
Statistical results	NG
<b>Conclusion Remarks</b>	No evidence of mutagenicity was seen under the test condition.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
Remarks for Data Reliability	Study was published in a peer review journal.
References	Sekizawa, J. and Shibamoto, T. (1982). Genotoxicity of Safrole-Related Chemicals in Microbial Test Systems. Mutation Research 101: 127-140.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	The DNA-repair test with Bacillus subtilis was performed as described by Kada et al. (1980).
Test Type	DNA-Repair test
System of Testing	Bacterial
GLP	NG
Year	1986

Species/Strain Bacillus subtilis/H17 Rec+ or M45 Rec-

**Metabolic Activation** S9 fraction was prepared from the PCB-treated male Sprague-

Dawley rats.

0.2 mg/disk **Doses/Concentration** 

**Statistical Methods** NG

Results No mutagenic effects in the absence of S9 fraction. DNA-repair

tests with S9 were not successful.

Genotoxic effects None

Statistical results NG

**Conclusion Remarks** No evidence of mutagenicity was detected under the test

conditions.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability Study was published in a peer review journal.

References Sekizawa, J. and Shibamoto, T. (1982). Genotoxicity of Safrole-

Related Chemicals in Microbial Test Systems. Mutation

Research 101: 127-140.

Substance Name	alpha-Amylcinnamaldehyde	
CAS No.	122-40-7	

122-40-7

Ames test was performed on five tester strains of Salmonella Method/guideline

typhimurium (TA 1535, TA 100, TA 1537, TA 1538, TA 98.

Reverse Mutation **Test Type** 

**Bacterial** System of Testing

**GLP** NG

Year 1983

Salmonella typhimurium (TA 1535, TA 100, TA 1537, TA 1538, Species/Strain

**Metabolic Activation** S-9 liver fraction was prepared from Aroclor-pretreated rats

(Aroclor 1254, 500 mg/kg, ip).

**Doses/Concentration** up to 3600 ug/plate

Statistical Methods Statistical significance was determined according to the

methods of Kastenbaum and Bowman (1970).

Positive controls were run in each experiment with the **Remarks for Test Conditions** 

reference mutagens sodium azide and benzo[a]pyrene.

No mutagenic activity was detected with any of the Salmonella Results

strains tested.

Cytotoxic concentration NG

**Genotoxic effects** None

**Conclusion Remarks** No mutagenic activity was detected with any of the Salmonella

strains tested.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Wild, D., King, M.-T., Gocke, E. and Eckhardt. (1983). Study of

> Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus Tests. Fd.

Chem. Toxicol. 21(6): 707-719.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	The antimutagenic effect of Cinnamaldehyde (CA) on the induction of HGPRT- mutants by methyl methanesulfonate (MMS), N-nitroso-N-methylurea (MNU), ethyl methanesulfonate (EMS) and UV light was investigated in the Chinese hamster V79 cell line.
Test Type	HGPRT- Mutants
System of Testing	Cell line
GLP	NG

Year 1994

Species/Strain Chinese hamster/V79 cell line

Metabolic Activation None

Doses/Concentration 50 or 100 mM.

**Statistical Methods** Student t-test

**Remarks for Test Conditions** Cells were seeded & then treated with UV light (12 J/m3) or

> MMS (2 mM), EMS (30 mM) or MNU (1 mM) for 1 h. Then the cells were washed 2X & the incubation was continued with fresh medium containing CA (0, 50 or 100 mM) for 2 or 4 h. Cell was washed, trypsinized & were seeded. The survival was measured by seeding 10E2 cells in a fresh medium. Mutation frequency was calculated as mutants/10E6 viable cells.

No mutagenic effect of CA; did not modify the mutation frequency when given to cells simultaneously with chemical mutagens MNU, EMS. MMS or UV; increased the cytotoxicity of

MMS but not of MNU & EMS

150 uM Cytotoxic concentration

Results

**Genotoxic effects** None

**Conclusion Remarks** Cinnamaldehyde inhibits some cellular function(s) promoting

> the repair of a variety of different cytotoxic lesions. At the same time, stimulation by Cinnamaldehyde of an error-free DNA repair mechanism acting on methyl methanesulfonate induced

mutagenic damage was indicated.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions. Remarks for Data Reliability Study was published in a peer review journal.

**References** Fiorio, R. and Bronzetti, G.(1994). Effects of Cinnamaldehyde

on Survival and Formation of HGPRT- Mutants in V79 Cells

Treated with Methyl Methanesulfonate, N-Nitroso-N-

Methylurea, Ethyl Methanesulfonate and UV Light. Mutation

Research 324: 51-57.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2

Method/guideline The chemical was tested in Strains of Salmonella using a liquid

preincubation procedure.

**Test Type** Reverse Mutation

System of Testing Bacterial

**GLP** NG

**Year** 1985

Species/Strain Salmonella typhimurium/TA104 & TA102

Metabolic Activation None

**Doses/Concentration** Tested up to the toxic concentration. (Unspecified)

Remarks for Test Conditions Use of two strains, TA104 and TA102 is described.

**Results** Negative

**Genotoxic effects** No mutagenic activity was reported.

**Conclusion Remarks**No mutagenic activity of Cinnamaldehyde was detected by the

use of two new base substitution strains TA104 and TA102.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Marnett, L.J., Hurd, H.K., Hollstein, M.C., Levin, D.E.,

esterbauer, H., and Ames, B.N. (1985). Naturally Occurring Carbonyl Compounds are Mutagens in Salmonella Tester

Strain TA104. Mutation Research 148: 25-34.

Substance Name	trans-Cinnamaldehyde (>95% pure)
CAS No.	104-55-2
Method/guideline	Cinnamaldehyde was tested for mutagenicity in five strains of Salmonella typhimurium both in the presence or absence of S-9 mix. Both the plate incorporation tests and the liquid preincubation assay were performed.

**Test Type** Reverse Mutation

System of Testing Bacterial

GLP NG

**Year** 1980

Species/Strain Salmonella typhimurium/TA1535, TA1537, TA1538, TA98 and

TA100

Metabolic Activation Rat or hamster liver homogenates from animals stimulated with

Aroclor 1254 (500 mg/kg 5 days).

**Doses/Concentration** 1- 500ug/plate

Results Negative

Genotoxic effects None

Conclusion Remarks No mutagenic activity of cinnamaldehyde was detected either

by the plate incorporation test or by the liquid preincubation assay in the presence or absence of rat or hamster S-9 fraction.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Lijinsky, W. and Andrews A.W. (1980). Mutagenicity of Vinyl

Compounds in Salmonella Typhimurium. Teratogenesis,

Carcinogenesis and Mutagenesis 1: 259-269.

Substance Name	Cinnamaldehyde	
CAS No.	104-55-2	

Method/guideline Reverse mutation assay using Salmonella typhimurium strains

TA92, TA1535, TA100, TA1537, TA94 and TA98 was carried

out according to the method of Ames.

**Test Type** Reverse Mutation

System of Testing Bacterial

**GLP** NG

**Year** 1984

Species/Strain Salmonella typhimurium/TA92, TA1535, TA100, TA1537, TA94

and TA98

Metabolic Activation Liver microsome fraction (S-9) prepared from the liver of

Fischer rats pretreated 5 days before with polychlorinated

biphenyls (500 mg/kg, ip).

Remarks for Test Conditions Solvent used DMSO

Results Cinnamic aldehyde induced 222 revertants (146 in control) at

0.5 mg/plate and 318 revertants (139 in control) at 0.1 mg/plate

in TA100 with and without S-9 mix, respectively.

Genotoxic effects Positive

Conclusion Remarks Cinnamic aldehyde was reported to be mutagenic in Salmonella

typhimurium strain TA100 in the presence and absence of S-9

mix.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

**References** Ishidate, M., Sofuni, T., Yoshikawa, K., Hauashi, M., Nohmi, T.,

Sawada, M. and Matsuoka. (1984). Primary Mutagenicity Screening of Food Additives Currently Used in Japan. Fd.

Chem. Toxic. 22(8) 623-636.

Substance Name	alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Method/guideline Ames test was performed on two tester strains of Salmonella

typhimurium TA 97and TA 102.

Test Type Reverse Mutation

System of Testing Bacterial

**GLP** NG

**Year** 1987

Species/Strain Salmonella typhimurium/TA97 and TA 102

Metabolic Activation S-9 liver fraction was prepared from Aroclor-pretreated rats

(Aroclor 1254, 500 mg/kg, ip).

**Doses/Concentration** 1-1000 ug/plate

Statistical Methods Kruskal-Wallis test

Remarks for Test Conditions Preincubation method using positive controls of 9-AA (20

ug/plate) for TA 97 with activation and 5 ug/plate without activation (S-9). Positive control for TA 102 was MMC (0.5 ug/plate) without activation and 9-AA (5 ug/plate) without

activation.

**Results** No mutagenic effects with or without S9 activation

Genotoxic effects None

Conclusion Remarks No mutagenic activity was detected with any of the Salmonella

strains tested.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Fujita H. and Sasaki M (1987) Mutagenicity Test of food

additives with Salmonella Typhirium TA 97 and TA102. Annals

of Tokyo Metr. Research Laboratory P.H. 38:423-430.

Substance Name alpha-Hexylcinnamaldehyde
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**CAS No.** 101-86-0

Method/guideline Ames test was performed on five tester strains of Salmonella

typhimurium (TA 1535, TA 100, TA 1537, TA 1538, TA 98.

**Test Type** Reverse Mutation

System of Testing Bacterial

**GLP** NG

**Year** 1983

Species/Strain Salmonella typhimurium (TA 1535, TA 100, TA 1537, TA 1538,

TA 98.

Metabolic Activation S-9 liver fraction was prepared from Aroclor-pretreated rats

(Aroclor 1254, 500 mg/kg, ip).

**Doses/Concentration** up to 3600 ug/plate

Statistical Methods Statistical significance was determined according to the

methods of Kastenbaum and Bowman (1970).

**Remarks for Test Conditions** Positive controls were run in each experiment with the

reference mutagens sodium azide and benzo[a]pyrene.

Results No mutagenic activity was detected with any of the Salmonella

strains tested.

Cytotoxic concentration NG

Genotoxic effects None

**Conclusion Remarks**No mutagenic activity was detected with any of the Salmonella

strains tested.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

References Wild D., King, M.T., Gocke, E. and Eckhardt. (1983). Study of

Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus tests Food

and Chemical Toxicology 21(6), 707-719.

Substance Name p-tert-Butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

Method/guideline Ames test was performed on five tester strains of Salmonella

typhimurium (TA 1535, TA 100, TA 1537, TA 1538, TA 98.

Test Type Reverse Mutation

System of Testing Bacterial

GLP NG

**Year** 1984

Species/Strain Salmonella typhimurium/TA 1535, TA 100, TA 1537, TA 1538,

TA 98

Metabolic Activation 4 or 10% Aroclor-induced S9 fraction prepared from the PCB-

treated male Sprague-Dawley rats

**Doses/Concentration** 0.0078 to 0.125 ul/plate

Statistical Methods NG

Remarks for Test Conditions Solvent, Ethanol. Plate incorporation method using positive

controls of 2-acetylaminofluorene (2ug/plate) for TA 98 and TA1538, mitomycin C (1 ug/plate) for TA102, so with activation

and 5 ug/plate without activation (S-9). Positive control.

Results No mutagenic activity was detected with any of the Salmonella

strains tested with or without S9 activation.

Cytotoxic concentration NG

Genotoxic effects None

Conclusion Remarks No mutagenic activity was detected with any of the Salmonella

strains tested.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Basic data given comparable to guidelines/standards.

**References** Givaudan-Roure (1984) Mutagenicity evaluation of *p-t-butyl-*

alpha-methylhydrocinnamaldehyde in the

Salmonella/mammalian plate incorporation assay. Unpublished

Report to RIFM.

Substance Name	<i>p</i> -tert-Butyl- <i>alpha</i> -methyldihydrocinnamaldehyde
CAS No.	122-40-7

Method/guideline Ames test was performed on five tester strains of Salmonella

typhimurium (TA 1535, TA 100, TA 1537, TA 98).

**Test Type** Reverse Mutation

System of Testing Bacterial

GLP NG

**Year** 1991

Species/Strain Salmonella typhimurium/TA 1535, TA 100, TA 1537, TA 98.

Metabolic Activation S9 fraction was prepared from the PCB-treated male Sprague-

Dawley rats.

**Doses/Concentration** 2.5 to 750 ug/plate without activation and 250 ug/plate with

activation.

Statistical Methods NG

Remarks for Test Conditions Solvent, DMSO.

Results No mutagenic activity was detected with any of the Salmonella

strains tested with or without S9 activation.

**Cytotoxic concentration** 667 ug/plate with (+S9), 333 ug/plate (-S9)

Genotoxic effects None
Statistical results NG

Conclusion Remarks No mutagenic activity

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Basic data given comparable to guidelines/standards.

Wagner V.O., and Twarszik, S. C. (1999) Bacterial reverse References

mutation assay of *p*-t-butyl-*alpha*-methyldihydrocinnamic

aldehyde. Unpublished journal.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline Test Type	Chromosomal aberration test was carried out using a Chinese hamster fibroblast cell line, CHL. The cells were exposed to 3 different doses for 24 and 48 hours. No metabolic activation system was applied.  Chromosomal aberration test
System of Testing	Chinese hamster fibroblast cell line CHL.
GLP	NG
Year	1984
Species/Strain	Chinese hamster fibroblast cell line CHL.
Metabolic Activation	None
Doses/Concentration	Max. dose = 0.015 mg/ml
Remarks for Test Conditions  Results	For quantitative evaluation of the clastogenic potential, the D20 was calculated, which is the dose (mg/ml) at which structural aberrations (including gaps) were detected in 20% of the metaphases observed. In addition, the TR value was calculated, which indicates the frequency of cells with exchange-type aberrations per unit dose (mg/ml). TR values are relatively high for chemicals that show carcinogenic potential in animals.  Cinnamic aldehyde was positive in chromosomal aberration test. TR value was 2133 and D20 = 0.01. It was also positive at 0.01 mg/ml at 24 h (20.0%, total incidence of cells with
Genotoxic effects	aberrations) and at 48 hr (15%, total incidence of cells with aberrations). The results were considered positive if the total incidence of cells with aberrations was 10.0% or more. Positive
Genotoxic effects Conclusion Remarks	aberrations) and at 48 hr (15%, total incidence of cells with aberrations). The results were considered positive if the total incidence of cells with aberrations was 10.0% or more. Positive  Cinnamic aldehyde was shown to be positive in chromosomal
	aberrations) and at 48 hr (15%, total incidence of cells with aberrations). The results were considered positive if the total incidence of cells with aberrations was 10.0% or more. Positive

Ishidate, M., Sofuni, T., Yoshikawa, K., Hauashi, M., Nohmi, T., Sawada, M. and Matsuoka. (1984). Primary Mutagenicity Screening of Food Additives Currently Used in Japan. Fd.

Chem. Toxic. 22(8) 623-636.

References

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline Test Type	The test chemical was screened for mutagenic activity using Salmonella typhimurium strains TA97, TA98, and TA100 with and without S9 metabolic activation using prolonged, non-standard preincubation time of up to 120 minutes. Reverse Mutation
System of Testing	Bacterial
GLP	NG
Year	1995
Species/Strain	Salmonella typhimurium Strain TA97, TA98 and TA100
Metabolic Activation	S9 fraction used but source not specified
Results	No mutagenic activity was detected
Genotoxic effects	None
Statistical results	NG
Data Qualities Reliabilities	Data appears to be reliable.
Remarks for Data Reliability	Reliability code 2. Reliable with restrictions.
References	Azizian, A. and Blevins, R.D. (1995). Mutagenicity and Antimutagenicity Testing of Six Chemicals Associated with the Pungent Properties of Specific Spices as Revealed by the Ames Salmonella/Microsomal Assay, Arch. Environ. Contam. Toxicol. 28: 248-258.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	The genotoxicity of cinnamaldehyde was studied by a bacterial mutation test in the Salmonella/microsome system with and without rat-liver microsome as the metabolic activation system.
Test Type	Reverse Mutation
System of Testing	Bacterial
GLP	NG
Year	1982
Species/Strain	Salmonella typhimurium Strain TA98 and TA100
Metabolic Activation	Rat-liver microsomes
Doses/Concentration	0.05 to 500 ug/plate

Results	Test substance did not induce a number of revertants that was
	over half of the number of spontaneous revertants of TA98 or
	TA100 either with or without S9 mix. Considerable mutagenic
	and the first common and the control of the control

activity was seen in positive standard mutagens.

Genotoxic effects None

**Conclusion Remarks**Cinnamaldehyde did not induce a number of revertants that

was over half of the number of spontaneous revertants of TA98

or TA100 either with or without \$9 mix.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Study was published in a peer review journal.

**References** Kasamaki, A., Takahashi, H, Niwa, J., Fujita, T. and Urasawa,

S. (1982). Genotoxicity of Flavoring Agents. Mutation Research

105: 387-392.

	100. 007 002.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Cinnamaldehyde was tested for genotoxicity using CH cell line B241 in culture stages between the 5th and 8th passages.
Test Type	Chromosomal aberration
System of Testing	Cell line
GLP	NG
Year	1982
Species/Strain	Chinese Hamster cell line B241
Metabolic Activation	Rat liver microsome from Sprague-Dawley rats treated with Aroclor 1254
Remarks for Test Conditions	One day after seeding, exponentially growing cells were exposed to the test chemical for 24 hrs, then incubated for another 24 hrs without chemical followed by treatment with colchicine (1 X 10-7M) for 2-3 hrs. Chromosome samples were prepared by the Giemsa staining method. Control cell cultures were treated with a medium containing DMSO equal in its concentration to the test solution of test chemical. The

Cinnamaldehyde induced severe chromosome aberration in the

treated CH cells suggesting a potential genotoxicity.

percentage of chromosome aberration was computed by scoring about 200 metaphase spreads, each containing 20-26

**Genotoxic effects** Induced severe chromosome aberration

Results

chromosomes.

**Remarks for Results** Various types of aberrations were observed in the treated cells,

such as severe chromatid break, chromosome break, chromatid

exchange, ring or dicentric chromosomes, fragmentation,

translocation and pulverization.

Conclusion Remarks Cinnamaldehyde induced severe chromosome aberration in the

treated CH cells suggesting a potential genotoxicity.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** The study was published in a peer review journal.

Kasamaki, A., Takahashi, H, Niwa, J., Fujita, T. and Urasawa, S. (1982). Genotoxicity of Flavoring Agents. Mutation Research References

105: 387-392.

## 4.3 In Vivo Genotoxicity

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Mouse bone marrow micronucleus assay.
Test Type	Micronucleus Test
GLP	NG
Year	1984
Species/Strain	Mice/ddY
Sex	Male
Route of administration	Intraperitoneal
Doses/concentrations	125, 250, 500 & 1000 mg/kg
Exposure period	18, 24, 30, 48, or 72 hrs
Remarks for Test Condition	Mice received one of the 4 different doses of the test material by IP and were killed after a time interval of 18, 24, 30, 48 or 72 hr following injection. Femoral marrow cells were smeared, fixed and stained. 100 polychromatic erythrocytes were scored and the number of micronucleated polychromatic erythrocytes were recorded.
Genotoxic effects	Not genotoxic
Remarks for Results	Micronucleated polychromatic erythrocytes did not increase in any dose or any sampling time. At 500 mg/kg more than 1 animal died; at 1000 mg/kg all animals died.
Conclusion Remarks	No evidence of genotoxicity.
Data Qualities Reliabilities	Reliability code 1. Reliable without restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.
References	Hayashi, M., Sofuni, T and Ishidate, M. (1984). A Pilot Experiment for the Micronucleus Test. The multi-sampling at multi-dose levels method. Mutation Research 141: 165-169.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Mouse
Test Type	Micronucleus test
GLP	NG
Year	1990
Species/Strain	ddY mice
Sex	Male
Route of administration	Oral
Doses/concentrations	250, 313, or 500 mg/kg bw
Exposure period	Single dose
Remarks for Test Condition  Genotoxic effects	Male ddY mice were irradiated with X-ray at 200 rad. After irradiation cinnamaldehyde was administered orally at 250, 313 or 500 mg/kg. In a time course study 500 mg/kg was given to mice immediately after the irradiation and the bone-marrow cells were sampled periodically. The micronucleus assay was performed according to the method described by Schmid 1976. Not genotoxic
Appropriate statistical evaluations?	Student t-test
Remarks for Results	A dose-dependent decrease in micronucleated polychromatic erythrocytes. At 500 mg/kg, there was 58% decrease in MNPCE. The test material did not increase the frequency of polychromatic erythrocytes, indicating that observed reduction of MNPCE was not a reflection of toxic effect of cinnamaldehyde on the bone-marrow.
Conclusion Remarks	Cinnamaldehyde reduced the frequency if X-ray induced micronuclei with toxicity of the test chemical to the bone marrow.
Data Qualities Reliabilities	Reliability code 1. Reliable without restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.
References	Sakasi, Y. F., Ohta, T., Imanishi, H, Watanabe, M., Matsumoto, K., Kato, T., and Shirasu, Y. (1990). Suppressing Effects of Vanilin, Cinnamaldehyde, and Anisaldehyde on Chromosome aberrations induced by X-rays in mice. Mutation Research 243: 299-302.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	DNA Fragmentation/Alkaline Elution Assay

**Test Type** DNA Fragmentation/Alkaline Elution Assay

**GLP** NG

**Year** 1994

Species/Strain Sprague-Dawley rats

**Sec** Male

Route of administration Oral

**Doses/concentrations** 1100 mg/kg

**Exposure period** Single dose. Animals killed after 3 hrs.

Remarks for Test Condition Male albino Sprague-Dawley rats were given by gastric

intubation a single dose (1100 mg/kg) of Cinnamaldehyde in carboxymethylcellulose. Rats were killed 3 hrs after treatment. DNA fragmentation (Single Strand break &/or Alkali-liable sites) was evaluated by the Alkaline Elution Technique.

Genotoxic effects None

Remarks for Results Cinnamaldehyde did not induce DNA fragmentation in liver

and gastric mucosa as evaluated by the alkaline elution

technique.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** The study was published in a peer-reviewed journal.

References Mereto, E., Brambilla-Campart, G., Ghia, M., Martelli, A., and

Brambilla, G. (1994), Cinnamaldehyde-induced micronuclei in

rodent liver. Mutation Research 322: 1-8.

Substance Name Cinnamaldehyde

**CAS No.** 104-55-2

Method/guideline Cinnamaldehyde induced micronuclei in rodent liver was

investigated.

**Test Type** Micronuclei Assay

**GLP** NG

**Year** NG

**Species/Strain** Male Albino Sprague-Dawley rats

**Sex** Male

Route of administration Oral

**Doses/concentrations** 550, 1100 or 1650 mg/kg.

**Exposure period** Single oral dose

Remarks for Test Condition Animals were subjected a 2/3 hepatectomy 20 hrs before

dosing in order to determine the clastogenic effect on hepatocytes and were killed 48 hrs after cinnamaldehyde administration. The frequency of micronucleated polychromatic erythrocytes was evaluated in marrow, hepatocytes and gastric

mucosa.

Genotoxic effects Not genotoxic

Remarks for Results No increase in the frequency of MNPCE in bone marrow 48 hrs

after administration of cinnamaldehyde; it induced a dosedependent clastogenic effect in the liver; significantly higher incidence of total nuclear anomalies but not of micronucleated

cells in forestomach mucosa

**Conclusion Remarks** High doses of cinnamaldehyde may produce a clastogenic

effect.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** The study was published in a peer-reviewed journal.

References Mereto, E., Brambilla-Campart, G., Ghia, M., Martelli, A., and

Brambilla, G. (1994). Cinnamaldehyde-induced micronuclei in

rodent liver. Mutation Research 322: 1-8.

Substance Name	Cinnamaldehyde

**CAS No.** 104-55-2

Method/guideline Cinnamaldehyde induced micronuclei in rodent liver was

investigated.

Test Type Micronuclei Assay

**GLP** NG

**Year** 1994

Species/Strain Male Albino Swiss mice

**Sex** Male

Route of administration Oral

**Doses/concentrations** 850, 1200 or 2550 mg/kg

**Exposure period** Single oral dose

Remarks for Test Condition Animals were subjected to a 2/3 hepatectomy 20 hrs before

dosing in order to determine the clastogenic effect on hepatocytes and were killed 48 hrs after cinnamaldehyde administration. The frequency of micronucleated polychromatic erythrocytes was evaluated in marrow, hepatocytes and gastric

mucosa.

Genotoxic effects Not genotoxic

Remarks for Results No increase in the frequency of MNPCE in bone marrow 48 hrs

after administration of cinnamaldehyde; it induced a dose-

dependent clastogenic effect in the liver.

Conclusion Remarks	High doses of cinnamaldehyde may produce a clastogenic.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.
References	Mereto, E., Brambilla-Campart, G., Ghia, M., Martelli, A., and Brambilla, G. (1994). Cinnamaldehyde-induced micronuclei in rodent liver. Mutation Research 322: 1-8.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Solt-Farber Assay System
Test Type	Solt-Farber Assay System
GLP	NG
Year	1994
Species/Strain	Male Sprague-Dawley rat
Sex	Male
Route of administration	Gavage
Doses/concentrations	500 mg/kg
Remarks for Test Condition  Genotoxic effects	Three groups of rats were initiated with NDEA (200 mg/kg ip). Two weeks later, Group 1: received 14 successive day of cinnamaldehyde; Group 2: rats were fed diet containing 0.02% 2 AAF (+ve control); Group 3: no treatment (-ve control). On day 7, all rats were hepatectomized. On day 28 all rats were killed and liver removed.  Not genotoxic
Remarks for Results  Conclusion Remarks  Data Qualities Reliabilities	Rats initiated with NDEA, administration of cinnamaldehyde for 14 days produce significant increase in average diameter & area of gamma-glutamyltraspeptidase positive foci that might be considered as indication of a potential promoting activity. The high doses of cinnamaldehyde may possibly a produce promoting effect in the liver of previously initiated animals. Reliability code 2. Reliable with restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.
References	Mereto, E., Brambilla-Campart, G., Ghia, M., Martelli, A., and Brambilla, G. (1994). Cinnamaldehyde-induced micronuclei in rodent liver. Mutation Research 322: 1-8.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2

Method/guideline DNA-Repair assay was carried out to examine the ability of the

test chemical to induce unscheduled DNA synthesis (UDS) or S-phase-synthesis (SPS) in Fischer-344 rats. Animals were administered the test chemical by oral gavage as a single bolus

DNA repair assay Test Type

**GLP** NG

Year 1989

Species/Strain Fischer-344 rats

Sex Male

Route of administration Oral

UDS: - 50, 200, or 1000 mg/kg. **Doses/concentrations** 

**Remarks for Test Condition** Doses were selected based approximately on the oral LD50

value and was selected as 80%, 40% and 10% of the LD50. Two doses were selected for SPS studies and three doses were utilized for UDS studies. SPS was examined at 48 hr post

treatment.

Cinnamaldehyde failed to induced UDS or SPS in rats at the Remarks for Results

doses tested.

**Conclusion Remarks** Cinnamaldehyde failed to induce the UDS or SPS in rats.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

The study was published in a peer-reviewed journal. Remarks for Data Reliability

References Mirsalis, J.C., Tyson, C.K., Steinmetz, K. L., Loh, E.K.,

> Hamilton, C.M., Bakke, J.P. and Spalding, J.W. (1989). Measurement of Unscheduled DNA Synthesis and S-Phase Synthesis in Rodent Hepatocytes Following In Vivo Treatment: Testing of 24 Compounds. Environmental and Molecular

Mutagenesis 14: 155-164.

**Substance Name** Cinnamaldehyde CAS No.

104-55-2

Method/guideline DNA

DNA repair assay **Test Type** 

**GLP** NG

1989 Year

Species/Strain B6C3F1 mice

Sex Male and Female

Route of administration Oral **Doses/concentrations** UDS: 50, 200, or 1000 mg/kg.

**Remarks for Test Condition** Doses selected based approximately on the oral LD50 value

> and selected as 80%, 40% and 10% of the LD50. Two doses were selected for SPS studies and three doses were utilized for

UDS studies. SPS was examined at 48 hr post treatment.

Genotoxic effects Not genotoxic

Cinnamaldehyde failed to induce UDS or SPS in mice at the Remarks for Results

doses tested.

Conclusion Remarks Cinnamaldehyde failed to induce the UDS or SPS in mice.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal.

References Mirsalis, J.C., Tyson, C.K., Steinmetz, K. L., Loh, E.K.,

Hamilton, C.M., Bakke, J.P. and Spalding, J.W. (1989). Measurement of Unscheduled DNA Synthesis and S-Phase Synthesis in Rodent Hepatocytes Following In Vivo Treatment: Testing of 24 Compounds. Environmental and Molecular

Mutagenesis 14: 155-164.

**Substance Name** p-tert-Butyl-alpha-methyldihydrocinnamaldehyde

CAS No. 80-54-6

Method/guideline Micronucleus test

**Test Type** Micronucleus test

**GLP** NG

Year 2000

Species/Strain ICR mice

Male and Female Sex

Route of administration Intraperitoneal

**Doses/concentrations** 150, 300, or 600 mg/kg

**Exposure** period Single intraperitoneal dose

**Remarks for Test Condition** Mice received one of the 3 different doses of the test material

> by IP and were killed after a time intervals of 48 or 72 hr following injection. Femoral marrow cells were smeared, fixed and stained with May-Gruenwald-Giemsa. 2000 polychromatic erythrocytes were scored and the number of micronucleated

polychromatic erythrocytes were recorded.

A slight increase (9/1000), males at 600mg/kg Genotoxic effects

NOEL (C)/LOEL (C) 300 mg/kg

Remarks for Results The authors noted the response was not biologically significant

since only one animal in the 600 mg/kg level had 3MNPCE

which is within the historical control range (0-7 MN/2000 PCE/animal. No significant increase and no dose-related increase was observed in any other group regardless of dose,

sex, or collection time.

Conclusion Remarks No evidence of genotoxicity

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**References** Gudi R. and Krsmanovic L. (2000) Mammalian erythrocyte

micronucleus test of para-tert-butyl-alpha-methylhydrocinnamic

aldehyde. Unpublished report.

Substance Name	alpha-Amylcinnamaldehyde

**CAS No.** 122-40-7

Method/guideline BASC test on Drosophila was performed as reported in

Eckhardt, King, Gocke and Wild, 1980.

Test Type BASC test

**GLP** NG

**Year** 19983

Species/Strain Insect, Drosophila melanogaster

**Sex** Male and Female

Route of administration Oral

**Doses/concentrations** 10 mM

**Remarks for Test Condition** The test substance to be fed to the flies was prepared in 5%

saccharose, with addition of 2% ethanol and 2% Tween 80. Ethyl nitrite was administered to Drosophila males in gaseous form. To do this flies were kept for 3 days in 1-liter bottle containing small amount of medium, and ethyl nitrite was

injected into the tightly closed bottles.

Genotoxic effects None

NOEL (C)/LOEL (C) 10 mM

Remarks for Results No mutagenic activity was demonstrated under the test

conditions

Conclusion Remarks No mutagenic activity was demonstrated under the test

conditions

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** The study was published in a peer-reviewed journal.

General Remarks A significant increase in sex-linked recessive lethal mutations

in single test but repeated tests did not confirm the mutagenic

activity. This anomaly was ascribed to chance.

References

Wild D., King, M.T., Gocke, E. and Eckhardt, K. (1983). Study of Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus Tests. Fd.

Chem. Toxic. 21(6): 707-719.

Micronucleus test. NMRI mice were treated once with the test material. The mice were killed and bone-marrow smear was prepared 30 hours after the treatment. The smears were
material. The mice were killed and bone-marrow smear was
stained according to the Schmid method & slides were scored.
Micronucleus test
NG
1983
NMRI mice
Male and Female
Not given
0, 405, 809, 1213 mg/kg
Dose: 0 mg/kg = 2.7 mean MNPE/1000PE; 405mg/kg=1.3 mean MNPE/1000 PE; 809 mg/kg=3.0 MNPE/1000 PE; 1213 mg/kg=1.5 MNPE/1000 PE PE = Polychromatic erythrocytes; MNPE = Micronucleated Polychromatic Erythrocytes. None
1213 mg/kg
No mutagenic activity was detected under the test conditions.
No mutagenic activity was detected under the test conditions.
Reliability code 1. Reliable without restrictions.
The study was published in a peer-reviewed journal.
Wild D., King, M.T., Gocke, E. and Eckhardt, K. (1983). Study of Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus Tests. Fd. Chem. Toxic. 21(6): 707-719.

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0

Method/guideline BASC test on Drosophila was performed as reported in

Eckhardt, King, Gocke and Wild, 1980.

Test Type BASC test

**GLP** NG

**Year** 19983

Species/Strain Insect, Drosophila melanogaster

**Sex** Male and Female

Route of administration Oral

**Doses/concentrations** 10 mM

Remarks for Test Condition The test substance to be fed to the flies was prepared in 5%

saccharose, with addition of 2% ethanol and 2% Tween 80. Ethyl nitrite was administered to Drosophila males in gaseous form. To do this flies were kept for 3 days in 1-liter bottle containing small amount of medium, and ethyl nitrite was

injected into the tightly closed bottles.

Genotoxic effects None

NOEL (C)/LOEL (C) 10 mM

Remarks for Results No mutagenic activity was demonstrated under the test

conditions. No of sex-linked lethals/chromosomes tested; Control: Brood I, 42/18.188; Brood II, 34/17,734; Brood III, 50/16,980 Test Material; Brood I, 10/2426; Brood II, 2/2418;

Brood III, 6/2405.

Conclusion Remarks No mutagenic activity was demonstrated under the test

conditions

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** The study was published in a peer-reviewed journal.

**General Remarks** A significant increase in sex-linked recessive lethal mutations

in single test but repeated tests did not confirm the mutagenic

activity. This anomaly was ascribed to chance.

**References** Wild D., King, M.T., Gocke, E. and Eckhardt, K. (1983). Study

of Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus Tests. Fd.

Chem. Toxic. 21(6): 707-719.

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Micronucleus test. NMRI mice were treated once with the test

material. The mice were killed and bone-marrow smear was

prepared 30 hours after the treatment. The smears were stained according to the Schmid method & slides were scored.

Test Type Micronucleus test

**GLP** NG

**Year** 1983

Species/Strain NMRI mice

Sex Male and Female

Route of administration Not given

**Doses/concentrations** 0, 324, 540, 756 mg/kg

Effect on mitotic index or PCE/NCE ration by dose

level and sex

Dose: 0 mg/kg = 1.0 mean MNPE/1000PE; 324mg/kg=2.1 mean MNPE/1000 PE; 540 mg/kg=1.8 MNPE/1000 PE; 756 mg/kg=2.4 MNPE/1000 PE

PE = Polychromatic erythrocytes;

MNPE = Micronucleated Polychromatic Erythrocytes.

Genotoxic effects None

**NOEL (C)/LOEL (C)** 756 mg/kg

**Remarks for Results**No mutagenic activity was detected under the test conditions.

**Conclusion Remarks** No mutagenic activity was detected under the test conditions.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**References** Wild D., King, M.T., Gocke, E. and Eckhardt, K. (1983). Study

of Artificial Flavouring Substances for Mutagenicity in the Salmonella/Microsome, BASC and Micronucleus Tests. Fd.

Chem. Toxic. 21(6): 707-719.

Substance Name Cinnamaldehyde

**CAS No.** 1104-55-2

Method/guideline Micronucleus test. 6C3F1 male and female mice

(10/group/sex) were maintained on diet with the test material for three months. Blood samples were taken and smears prepared. Frequency of micronuclei per 1000 polychromatic erythrocytes and 2000 normochromatic erythrocytes were

measured in up to 5 animals per group.

Test Type Micronucleus test

**GLP** NG

Year 2003

Species/Strain B6C3F1 mice

Sex Male and Female

Route of administration Dietary

**Doses/concentrations** 0, 4100,8200,16,500, and 33,000 ppm microencapsulated in the diet

Effect on mitotic index or Males

PCE/NCE ration by dose level and sex

Dose:0 ppm = 2.1 mean MNPE/1000PE; 4100 ppm=1.9 mean MNPE/1000 PE; 8200 ppm=1.9 MNPE/1000 PE; 16,500 ppm=2.0 MNPE/1000 PE

33,00 ppm=1.7 MNPE/1000 PE

**Feales** 

Dose:0 ppm = 1.9 mean MNPE/1000PE; 4100 ppm=1.5 mean MNPE/1000 PE; 8200 ppm=1.8 MNPE/1000 PE; 16,500 ppm=1.9 MNPE/1000 PE 33,00 ppm=1.4 MNPE/1000 PE

PE = Polychromatic erythrocytes;

MNPE = Micronucleated Polychromatic Erythrocytes.

Genotoxic effects None

**NOEL** 33,000ppm

**Remarks for Results**No mutagenic activity was detected under the test conditions.

**Conclusion Remarks**No mutagenic activity was detected under the test conditions.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** The study was published in a peer-reviewed journal.

References NTP (2003) National Toxicology Program. Toxicology and

carcinogenesis studies of trans-cinnamaldehyde

(microencapsulated) (CAS No. 14371-10-9) in F344/N rats and B6C3F1 Mice. (Feed studies)Technical Report Series 5014NIH Publication No. 02-4448. U.S. Department of Health and

**Human Services** 

## 4.4 Repeat Dose Toxicity

Substance Name	Cinnamaldehyde
CAS	104-55-2
Remarks for Substance	Assay: >95%
Method/guideline	National Toxicology Program 2-yr Bioassay
Year	2003
Species/Strain	Rat/F344/N
Sex	Male and Female
Route of administration	Oral-dietary (microencapsulated)

**Doses/concentration levels** 1,000, 2,100, or 4,100 ppm cinnamaldehyde microencapsulated

for 104 to 105 weeks

**Exposure period** 104 - 105 weeks

Frequency of treatment Continuous

**Control Group** Basal diet containing placebo microcapsules

Remarks for test conditions A carcinogenicity study was conducted in which groups of 50

F344/N rats of each sex were maintained on diets containing 0 (untreated control), 0 (vehicle control containing placebo microcapsules), 1,000, 2,100, or 4,100 ppm cinnamaldehyde microencapsulated for 104 to 105 weeks. The dietary concentrations were calculated to provide an average daily intake of 50, 100, or 200 mg/kg bw per day. Feed analysis for concentrations are as specified in the study with mice (see below). Female rats were housed five per cage, while male rats were housed two or three per cage. Feed and water were provided ad libitum. Clinical findings were performed twice daily and body weights were monitored on days 0 (body weights only), 8, and 36 and then every four weeks until the end of the study. Seven (7)-day feed consumption was measured every 4 weeks during the study. Gross and histopathological

examinations were performed on all animals at termination of

the study.

NOAEL(NOEL) 4100 ppm

Toxic response/effects by dose level

Mean body weights of high-dose females and males (4,100 ppm) were less than those of the vehicle control after weeks 25 and 49, respectively. Feed consumption in the 2100 and 4100 ppm group of males and 4100 ppm female group was less than that of the respective control group. Survival of all treated groups of male rats was greater than that for the vehicle control group. No neoplastic or non-neoplastic lesions were observed that could be associated with the administration of diets containing 1,000, 2,100, or 4,100 ppm of cinnamaldehyde.

Statistical evaluations Yes

**Remarks for results**Based on the above observations, the NTP concluded "under

the conditions of these 2-year feed studies there was no evidence of carcinogenic activity of cinnamaldehyde in male or

female rats exposed to 1,000, 2,100, or 4,100 ppm.

**Conclusion remarks** The results of the NTP study in both sexes of F344 rats and

B6C3F1 mice indicate that under conditions of the 2-year feed studies, cinnamaldehyde shows no evidence of carcinogenic

potential in rodents.

**Data Qualities Reliabilities** Reliability Code 1. Reliable without restriction.

**References** NTP (2003) National Toxicology Program. Toxicology and

carcinogenesis studies of trans-cinnamaldehyde

(microencapsulated) (CAS No. 14371-10-9) in F344/N rats and B6C3F1 Mice. (Feed studies)Technical Report Series 5014NIH

Publication No. 02-4448. U.S. Department of Health and Human Services, Public Health Service, National Institutes of Health.

Substance Name Cinnamaldehyde

CAS 104-55-2

Remarks for Substance Assay: >95%

Method/guideline National Toxicology Program 2-yr Bioassay

Year 2003

Species/Strain Mouse/B6C3F1

Sex Male and Female

Route of administration Oral-dietary (microencapsulated)

Doses/concentration levels 1,000, 2,100, or 4,100 ppm cinnamaldehyde microencapsulated

for 104 to 105 weeks

Exposure period 104 - 105 weeks

Frequency of treatment Continuous

Control Group Basal diet containing placebo microcapsules

Remarks for test conditions A carcinogenicity study was conducted in which groups of 50

B6C3F1 mice of each sex were maintained on diets containing 0 (untreated control), 0 (vehicle control containing placebo microcapsules), 1,000, 2,100, or 4,100 ppm cinnamaldehyde microencapsulated for 104 to 105 weeks. The dietary concentrations were calculated to provide an average daily intake of 125, 270, or 550 mg/kg bw per day. Feed analysis for concentrations are as specified in the study with rats. Female mice were housed five per cage, while male mice were housed singly per cage. Feed and water were provided ad libitum. Clinical findings were performed twice daily and body weights were monitored on days 0 (body weights only), 8, and 36 and then every four weeks until the end of the study. Seven (7)-day feed consumption was measured every 4 weeks during the study. Gross and histopathological examinations were performed on all animals at termination of the study.

NOAEL(NOEL) 4100 ppm

Toxic response/effects by dose

level

Mean body weights of high-dose females and males (4,100 ppm) were less than those of the vehicle control after weeks 25 and 49, respectively. Feed consumption in the 2100 and 4100 ppm group of males and 4100 ppm female group was less than that of the respective control group. Survival of the 2100 ppm group of males was less than that of the control group. No neoplastic or non-neoplastic lesions were observed that could be associated with the administration of diets containing 1,000,

2,100, or 4,100 ppm of cinnamaldehyde.

Statistical evaluations Yes

Remarks for results Based on the above observations, the NTP concluded "under

the conditions of these 2-year feed studies there was no evidence of carcinogenic activity of cinnamaldehyde in male or

female mice exposed to 1,000, 2,100, or 4,100 ppm.

Conclusion remarks The results of the NTP study in both sexes of F344 rats and

B6C3F1 mice indicate that under conditions of the 2-year feed studies, cinnamaldehyde shows no evidence of carcinogenic

potential in rodents.

Data Qualities Reliabilities Reliability Code 1. Reliable without restriction.

References NTP (2003) National Toxicology Program. Toxicology and

carcinogenesis studies of trans-cinnamaldehyde

(microencapsulated) (CAS No. 14371-10-9) in F344/N rats and B6C3F1 Mice. (Feed studies)Technical Report Series 5014NIH Publication No. 02-4448. U.S. Department of Health and Human Services, Public Health Service, National Institutes of

Health.

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

Method/guideline Test Material was administered orally for 13 weeks to 3 groups

of 6 beagle dogs by means of gelatin capsules. Six dogs were

kept as controls and received the empty gelatin capsules.

**GLP** Yes

**Year** 1990

Species/Strain Beagle dogs

Sex Male and Female

Route of administration Oral

**Doses/concentration** 4.4, 22.3 or 44.6 mg/kg

**Exposure** 91 days

Frequency of treatment Daily

**Control Group and treatment** Yes

Post exposure observation

period

NG

NOAEL (NOEL) 44.6 mg/kg

LOAEL (LOEL) No adverse effects at highest dose

Actual dose received by

dose level and sex

Toxic response/effects by

dose level

vei

Statistical evaluations DUNN test

**Remarks for Results**No adverse effect with respect to the general state of health,

the body weight development, the behavior of the dogs,

hematological & clinical chemical parameters & opthalmoscopy,

macroscopic, pathology & histological appearance of the

organs and tissues examined were noted.

Conclusion Remarks This study demonstrates a NOAEL in dogs of at least 44.6

mg/kg/day.

NG

None

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

**Remarks for Data Reliability** The study was conducted in accordance with GLP.

**References** Givaudan Roure (1990b). A toxicity study following oral

administration of *p*-t-butyl alpha-methylhydrocinnamic aldehyde in dogs during a period of 13 weeks. Unpublished Report to

RIFM.

**Substance Name** p-t-Butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Method/guideline** Test material was orally administered at 200 mg/kg/day to 3

female beagle dogs by means of gelatine capsules for a period of 91 days. Incompatibility reactions, body weights & the group

feed intake were recorded.

**GLP** NG

**Year** 1990

Species/Strain Beagle dogs

Sex Female

Route of administration Oral

**Doses/concentration** 200 mg/kg/day

**Exposure** 13 weeks

**Control Group and treatment** Yes

Post exposure observation

period

NG

Remarks

Blood chemistry tests & an autopsy were performed. Blood parameters measured: Aspartate aminotransferase,

cholinesterase, cholesterol, alkaline phosphatase, gamma-

alutamiltransferase.

NOAEL (NOEL) 200 mg/kg

Actual dose received by dose level and sex

NA

Toxic response/effects by

dose level

None

Statistical evaluations

NG

Remarks for Results The administration of test material was asymptomatically

tolerated. The development of the body weights were

unaffected by the intake of the test article. The feed intake was normal. No treatment related blood chemistry changes were seen; especially, no reduction of plasma cholinesterase occurred. There were significant findings at necropsy.

This study demonstrates a NOAEL in dogs of 200 mg/kg.

Conclusion Remarks This study demonstrates a NOAEL in dogs of 200 mg/kg

bw/day.

**Data Qualities Reliabilities** Reliability code1. Reliable without restrictions.

**References** Givaudan-Roure (1990f) A complementary oral toxicity study

with *p*-t-butyl *alpha*-methylhydrocinnamic aldehyde on female dogs during a period of 13 weeks. Unpublished Report to RIFM

**Substance Name** *p*-t-Butyl-*alpha*-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

Method/quideline Test material was administered orally for 9 weeks to 2 beagle

dogs by means of gelatine capsules. Six dogs were kept as

controls and received the empty gelatine capsules.

**GLP** NG

**Year** 1990

Species/Strain Beagle dogs

**Sex** Male

Route of administration Oral

**Doses/concentration** 50 ul/kg bw/day for days 1-7, 100 ul/kg bw/day for days 8-14,

200 ul/kg bw/day for days 15-21, 400 ul/kg bw/day for days 22-

50, 600 ul/kg bw/day for days 50-64.

**Exposure** 64 days

Frequency of treatment Daily

Control Group and treatment None

Post exposure observation

period

None

Remarks for Test Conditions Clinical signs and body weights were recorded daily and

hematological examinations and clinical chemistry

determinations were performed weekly. Histopathology of brain, spinal cord, sciatic nerve, ulnar nerve, liver, kidney, and testes

were performed at week 9.

NOAEL (NOEL) 400 ul/kg/day

LOAEL (LOEL) None

Actual dose received by dose level and sex

Dose of 400 ul/kg/day administered from days 22-50 of the

study.

Toxic response/effects by

dose level

None

Statistical evaluations

NG

One dog showed increased GPT from week 7 onward and Remarks for Results

> increased GLDH from week 4 onward. Mild changes in the seminiferous epithelium of both dogs were not significantly

different from that seen in untreated dogs.

Pilot study that did not establish evidence of testicular effects in **Conclusion Remarks** 

dogs.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability The study was conducted in accordance with GLP.

References Givaudan Corporation (1990e) Pilot study on male dogs with p-

t-butyl-alpha-methylhydrocinnamic aldehyde following oral

administration (increasing dosage) during 9 weeks.

Unpublished report to RIFM.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Weanling Osborn-Mendel rats were fed diet containing 1000,
GLP	2500, or 10,000 ppm of the test substance for 16 weeks. Pre GLP
Year	1967
Species/Strain	Osborne-Mendel rats
Sex	Male and Female
Doses/concentration	1000, 2500 and 10,000 ppm
Exposure	16 weeks
Frequency of treatment	Daily in the diet

Post exposure observation

Control Group and treatment

period

NG

NOAEL (NOEL)

2500 ppm

Actual dose received by

dose level and sex

NA

Statistical evaluations

NG

Remarks for Results No effects were seen at 1000 or 2500 ppm. At 10,000 ppm,

Diet containing corn oil

slight hepatic cell swelling and slight hyperkeratosis of

squamous portion of stomach was noted.

**Conclusion Remarks** NOAEL for cinnamaldehyde was shown to be 2500 ppm in rat

by oral route.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions. **Remarks for Data Reliability** This study was published in a peer-reviewed journal.

References Hagan, E.C., Hansen, W.H., Fitzhugh, O.G., Jenner, P.M.,

Jones, W.I., Taylor, J.M., Long, E.L., Nelson, A.A., and Brouwer, J.B. (1967). Food Flavourings and Compounds of Related Structure. II. Subacute and Chronic Toxicity. Fd.

Cosmet. Toxicol. 5: 141-157.

	Cosmet. Toxicol. 5: 141-157.
Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	10 rats were fed a diet containing cinnamaldehyde (est. daily intake 50, 100 & 200 mg/kg) for 12 wks. Physical appearance, behavior and efficiency of food utilization were calculated.
GLP	Pre GLP
Year	1958
Species/Strain	Rats
Sex	Male and Female
Route of administration	In diet
Doses/concentration	Estimated daily intake: 50, 100 or 200 mg/kg
Exposure	12 weeks
Frequency of treatment	Daily
Control Group and treatment	Yes
Post exposure observation	NG
period Remarks	After 12 days of treatment, urine of 3 male and 3 female rats were examined for sugar and albumin and blood hemoglobin levels were also determined.
NOAEL (NOEL)	200 mg/kg

Actual dose received by

dose level and sex

Toxic response/effects by

dose level

Statistical evaluations

**Remarks for Results** 

**Conclusion Remarks** 

**Data Qualities Reliabilities** 

**General Remarks** 

NA

None

NG

No statistically significantly differences were observed between treated and control groups. No adverse effects were observed

on growth, food intake, efficiency of food utilization or other physiological criteria.

NOAEL was determined to be 200 mg/kg.

Reliability code 1. Reliable without restrictions.

This is a follow-up study for Trubek Laboratories 1958a.

Trubek Laboratories (1958b). Toxicological Examination of Cinnamic Aldehyde (Class IV, Part 2). References

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Rats were fed test ration containing Cinnamic aldehyde (897ppm), methyl cinnamate (25ppm), ethyl cinnamate (25ppm), cinnamyl cinnamate (25ppm) and <i>alpha</i> methyl cinnamic aldehyde (25ppm) for 12 weeks. Autopsies were performed on all rats.
GLP	Pre GLP
Year	1958
Species/Strain	Rat
Sex	Male and Female
Route of administration	Diet
Doses/concentration	Cinnamic aldehyde (897ppm), methyl cinnamate (25ppm), ethyl cinnamate (25ppm), cinnamyl cinnamate (25ppm) and <i>alpha</i> methyl cinnamic aldehyde (25ppm)
Exposure	12 weeks
Frequency of treatment	Continuously in diet
Control Group and treatment	Yes
Post exposure observation period Remarks	After 12 weeks of treatment, urine from 3 males and 3 females were examined for presence of sugar and albumin and blood hemoglobin levels. Autopsies were performed on all rats. Body weights and organ weight were recorded
Actual dose received by	weights and organ weight were recorded. NG
dose level and sex Toxic response/effects by dose level	Growth of male rats was retarded but not statistically significant at p=0.05. Food intake was not adversely affected. Food intake was not adversely affected. Efficiency of food utilization for both sexes was significantly depressed (male p=0.01 & female p=0.05). Urine was free of sugar and albumin. Blood hemoglobin was normal.
Statistical evaluations	NG
Conclusion Remarks	The cinnamate mixture was shown to depress the efficiency of food utilization in both sexes.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
General Remarks	See Trubek 1958b for a follow-up study.
References	Trubek (1958a). Toxicological Screening of Components of Food Flavors. Class IV. Cinnamates

040 N	404.00.0	
Substance Name	alpha-Hexylcinnamaldehyde	

**CAS No.** 101-86-0

Method/guideline Test material was applied percutaneously to the shaved dorsa

of 10 male Sprague-Dawley rats at dose levels of 0.125, 0.25,

0.50 & 1.00 g/kg/day for 90 consecutive days,

**GLP** GLP

**Year** 1980

Species/Strain Sprague-Dawley rats

Sex Male and Female

Route of administration Percutaneous

**Doses/concentration** 0.125, 0.25, 0.50 & 1.00 g/kg/day

**Exposure** 90 days

Frequency of treatment Daily

Control Group and treatment Yes

**Remarks** Parameter monitored: Body wt., Food consumption,

Hematology, Ophthalmological examination, Blood chemistry (BUN), serum glutamic pyruvic transaminase, serum glutamic oxalacetic transaminase, total bilirubin, fasting serum glucose, serum alkaline phosphatase), urinalysis, Gross Pathology,

Histopathology.

LOAEL (LOEL) 0.125 g/kg/day

Toxic response/effects by dose level

Dose-dependent dermal irritation characterized by erythema, cracking, dryness & sloughing; 5 male & 3 female from 1.0 g/kg died before 90 days: increased food consumption in females @ 0.25, 0.50, & 1.00 g/kg; inconsistent changes in hemoglobin, hematocrit, erythrocyte count, SGOT & SGPT; consistent elevation in white blood cell and the segmented neutrophil counts @ 0.50 & 1.00 g/kg; reduced lymphocyte count in males @ 1.00 g/kg; elevated white blood cell count in females @ 0.25-1.00 g/kg; reduced serum glucose & increased BUN & SAP in all rats; dose-dependent irritation of the GI-tract and the treated skin; increased liver & kidney wt in female @ 0.25-1.00 a/kg; at 1.00 g/kg; hepatic hydropic vacuolization & single cell degeneration, splenic lymphoid depletion & fibrosis, focal gastric ulceration & chronic necrotizing dermatitis with acanthosis, hyperkeratosis & sebaceous gland hyperplasia; dose-dependent increases in the myeloid-erythroid &

decreases of the cell-fat ratios.

Conclusion Remarks Percutaneous administration of Hexyl Cinnamic Aldehyde for 90

days produced multisystemic toxicity in the rats.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** The study was conducted in accordance with GLP.

**References** Lough, R., Owston, E., Klein, G., Qureshi, S., and Bier, C.

(1980). A subacute (90 Day) Percutaeous Toxicity Study of Hexyl Cinnamic Aldehyde in the Albino Rat. Unpublished. Bio-Research Lab. Report to RIFM.

Substance Name	Cinnamalydehyde
CAS No.	104-55-2
Method/guideline	Subchronic study. 10 mg of test substance was given every other day in normal or low protein diet (9% casein). Duration not given.
GLP	No
Year	1965
Species/Strain	Not reported
Sex	Not reported
Route of administration	Diet
Doses/concentration	10 or 50 mg every other day
Exposure	Not reported
Frequency of treatment	Every other day
Control Group and treatment	Not reported
Post exposure observation period	NG
Remarks	Article in Romanian. No details given in the English Abstract.
LOAEL (LOEL)	10 mg
Actual dose received by dose level and sex	NG
Toxic response/effects by dose level	The activity of liver aldolase showed significant increase and the activity of succindehydrogenase showed a tendency to decrease.
Statistical evaluations	NG
Remarks for Results	No effect on weight gain, food ingestion and protein efficiency. No effect on the liver weight and ascorbic acid content and the aspartic glutamic transaminase activity
Conclusion Remarks	Administration of test substance (10 mg) resulted in increased activity of liver aldolase and the activity of succindehydrogenase showed a tendency to decrease.
Data Qualities Reliabilities	Reliability code. 3. Data not reliable.
Remarks for Data Reliability	Article in Romanian. No details given in the English Abstract.
References	Sporn A. (1965). Investigation of the Toxicity of Cynamic Aldehyde. Igiena 14(6): 339-346.

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	Two male rhesus monkeys (Macaca Mulatta) were orally administered with 100 mg/kg/day of test substance suspended in fluid-baby food for 5 consecutive days.
GLP	NG
Year	1990
Species/Strain	Rhesus monkey Macaca Mulatta
Sex	Male
Route of administration	Oral in food
Doses/concentration	100 mg/kg/day for 5 consecutive days.
Exposure	5 days
Frequency of treatment	Daily
Remarks	At the end of the study, the Rhesus monkey were anesthetized and perfused with glutaraldehyde. Testes and epididymies were microscopically examined.
NOAEL (NOEL)	100 mg/kg
Toxic response/effects by dose level	None
Remarks for Results	No changes in body weight or testes were noted.

References	Givaudan Roure (1990g). A 5-day oral toxicity study with <i>p</i> -t-

**Conclusion Remarks** 

**Data Qualities Reliabilities** 

butyl-*alpha*-methylhydrocinnamic aldehyde in male rhesus monkeys. Unpublished, Report to RIFM.

mg/kg for 5 days. Reliability code 2. Reliable with restrictions.

No toxic effects were observed in monkeys treated with 100

Substance Name	p-t-Butyl-alpha-methyldihydrocinnamaldehyde
CAS No.	80-54-6
Method/guideline	The test material was dermally administered to male albino rats at dose levels of 250, 500, 1000 & 2000 mg/kg/day for 5 days.
GLP	NG
Year	1991
Species/Strain	Albino rats
Sex	Male
Route of administration	Dermal

**Doses/concentration** 250, 500, 1000 and 2000 mg/kg/day for 5 days.

**Exposure** 5 days

Frequency of treatment Daily

Remarks The mortalities, adverse symptoms & lower body weights were

recorded. At termination, all rats were euthanized and subjected to a full necropsy. Testes and epididymides were

microscopically examined.

NOAEL (NOEL) 1000 mg/kg/day

LOAEL (LOEL) 2000 mg/kg/day

Toxic response/effects by

dose level

No chemical related mortalities, Initial disturbance of body weight at 2000 mg/kg. No compound related gross lesions;

Atrophy in the testes at 2000 mg/kg/day.

**Conclusion Remarks** Treatment of rat with 2000 mg/g/day for 5 days dermally

resulted in disturbance in body weight and atrophy in the testes.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Givaudan Roure (1991). A 5-day toxicity study with *p*-t-butyl-

alpha-methy-hydrocinnamic aldehyde on male rats: dermal administration compared to oral administration. Unpublished.

Report to RIFM.

**CAS No.** 80-54-6

Method/guideline The test material was orally administered to male albino rats at

dose levels of 25, 50 and 100 mg/kg/day for 5 days.

**GLP** NG

**Year** 1991

**Sex** Male

Route of administration Oral

**Doses/concentration** 25, 50 and 100 mg/kg/day for 5 days.

**Exposure** 5 days

Frequency of treatment Daily

**Remarks** The mortalities, general symptoms & body weights were

recorded. At termination, all rats were euthanized and subjected to a full necropsy. Testes and epididymides were

microscopically examined.

NOAEL (NOEL) 25 mg/kg/day

LOAEL (LOEL) 50 mg/kg/day

Toxic response/effects by

dose level

No chemical related mortalities, Initial disturbance of body weight at 50 and 100 mg/kg. No compound related gross

lesions; Atrophy in the testes at 50 and 100 mg/kg/day.

Conclusion Remarks Treatment of rat with 50 or 100 mg/g/day for 5 days orally

resulted in disturbance in body weight and atrophy in the testes.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Givaudan Roure (1991). A 5-day toxicity study with *p*-t-butyl-

alpha-methy-hydrocinnamic aldehyde on male rats: dermal administration compared to oral administration. Unpublished.

Report to RIFM.

**Substance Name** alpha-Amylcinnamaldehyde (at least 97% pure), pale yellow

liquid with a floral odor.

**CAS No.** 122-40-7

Method/guideline 15 Male & 15 female rats were fed diet containing the test

substance for 14 weeks at dietary levels of 0, 80, 400 or 4000 ppm. Rats were killed by exanguination under barbiturate anesthesia. Parameters monitored were: body wt, hemoglobin

content see below.

**GLP** NG

**Year** 1973

Species/Strain Rats CFE strain

Sex Male and Female

Route of administration Oral

**Doses/concentration** 0, 80, 400 or 4000 ppm

**Exposure** 14 weeks

Frequency of treatment Continuous

**Control Group and treatment** Diet without the test material

**Remarks** Parameters measured: packed cell volume, counts of

erythrocytes, total leucocytes & individual types of lecocytes, serum, urea, glucose, total protein, albumin, activation of glutamic oxaloacetate & glutamic-pyruvic transaminase & lactic dehydrogenase, urinalysis for the final week of treatment. Each

animal was given an autopsy.

NOAEL (NOEL) 400 ppm

LOAEL (LOEL) 4000 ppm

Actual dose received by dose level and sex

Toxic response/effects by

**Remarks for Results** 

dose level

Male: 6.1, 29.9 or 287.3 mg/kg/day; female:6.7, 34.9 or 320.3

mg/kg/day

Increase in the relative liver & kidney weights of the rats fed diet

containing 4000 ppm of the test substance for 14 weeks.

These were not associated with any histopathological changes. No differences over controls were seen in the rate of body wt

gain, the consumption of food & water, hematological

measurements, serum analyses, urinary cell excretion or renal

concentration tests.

**Conclusion Remarks** NOAEL for the test material was shown to be 400 ppm.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal.

References Carpanini, F.M.B., Gaunt, I.F., Wright, M.G., Grasso, P. and

Gangolli, S.D. (1973), Short-Term Toxicity of Amyl Cinnamic

Aldehyde in Rats. Fd. Cosmet. Toxicol. 11: 725-734.

**Substance Name** Cinnamaldehyde (98% pure)

**CAS No.** 104-55-2

**Method/guideline** Rats were administered the test substance by gavage for 2 wks.

**GLP** NG

**Year** 1994

Species/Strain F344/N rats

**Sex** Male and Female

Route of administration Gavage

**Doses/concentration** 0, 235, 470, 940, 1880 & 3750 mg/kg/day for 14 days

Exposure 14 days

Frequency of treatment Daily

Control Group and treatment Corn oil gavage

**Remarks** A complete autopsy was performed on all animals that died,

and at the termination of the study on all treated and control animals. Body wt and the organ wt. of the liver, right kidney and

the spleen were determined.

NOAEL (NOEL) 235 mg/kg/day

LOAEL (LOEL) 470 mg/kg/day

Toxic response/effects by

dose level

All rats dosed at 1880 & 3750 mg/kg/day died or were killed when moribund during the first 7 days of dosing. Microscopic

lesions included a minimal to moderate forestomach

hyperplasia in males at doses of 470 mg/kg/day and higher.

Statistical evaluations ANOVA

Remarks for Results There were no consistent differences in organ wt or organ wt:

body wt ratios between surviving treated or controls. Clinical

signs and gross lesions were absent in surviving rats.

Conclusion Remarks Test substance at dose 470 mg/kg/day and above produces

forestomach hyperplasia and was lethal at dose of 1880 and

above.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal.

References Herbert, C. D., Yuan, J. and Dieter, M.P. (1994). Comparison of

the Toxicity of Cinnamaldehyde When Administered by Microencapsulation in Feed or by Corn Oil Gavage. Food and

Chemical Toxicology 32(12): 1107-1115.

Substance Name	Cinnamaldehyde (98% pure)

**CAS No.** 104-55-2

Method/guideline Mice were administered the test substance by gavage for 3

weeks

**GLP** NG

**Year** 1994

Species/Strain B6C3F1 mice

**Sex** Male and Female

Route of administration Gavage

**Doses/concentration** 656, 1310, 2620, 5250 & 10500 mg/kg/day

Exposure 21 days

Frequency of treatment Daily

Control Group and treatment Corn-oil gavage

**Remarks** A complete autopsy was performed on all animals that died,

and at the termination of the study on all treated and control animals. Body wt and the organ wt. of the liver, right kidney and

the spleen were determined.

NOAEL (NOEL) 656 mg/kg/day

LOAEL (LOEL) 1310 mg/kg/day

Toxic response/effects by

dose level

All mice gavaged at doses of 5250 and 10,500 mg/kg/day, as well as all female mice and three male mice dosed with 2620

mg/kg/day died within first 2 days. No clinical signs, or gross or microscopic lesions were visible in these mice. The only

microscopic lesions observed in surviving mice were a minimal

to mild forestomach hyperplasia & a minimal kidney nephropathy at doses of 1310 mg/kg/day and higher.

Statistical evaluations ANOVA

Conclusion Remarks Test substance at doses 1310 mg/kg/day and above produce

forestomach hyperplasia and was lethal at dose of 5250 and

above.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal.

References Herbert, C. D., Yuan, J. and Dieter, M.P. (1994). Comparison of

the Toxicity of Cinnamaldehyde When Administered by Microencapsulation in Feed or by Corn Oil Gavage. Food and

Chemical Toxicology 32(12): 1107-1115.

Substance Name	Cinnamaldehyde (98% pure)
CAS No.	104-55-2
Method/guideline	A group of rats were fed a diet containing 0, 0.625, 1.25, 2.5,
GLP	5.0 or 10% Cinnamaldehyde microcapsules for 14 days. NG
Year	1994
Species/Strain	F344/N rats
Sex	Male and Female
Route of administration	Oral in diet
Doses/concentration	A group of rats were fed a diet containing 0, 0.625, 1.25, 2.5,
Exposure	<ul><li>5.0 or 10% Cinnamaldehyde microcapsules for 14 days.</li><li>14 days</li></ul>
Frequency of treatment	Continuous
Remarks	A complete autopsy was performed on all animals that died, and at the termination of the study on all treated and control animals. Body wt and the organ wt. of the liver, right kidney and the spleen were determined.
NOAEL (NOEL)	0.625%
LOAEL (LOEL)	1.25%
Toxic response/effects by dose level  Statistical evaluations	Marked dose-related depression in body wt gain, slight decrease in spleen: body wt ratio for male rats in 10% group, dose dependent decrease in food consumption. Gross lesions in both sexes were limited to a reduction in the size of reproductive organs and secondary sex glands (seminal vesicles & prostates of males & ovaries & uteri of females). Hyperplasia of the forestomach ANOVA
Conclusion Remarks  Data Qualities Reliabilities	Treatment of rat with microencapsulated cinnamaldehyde resulted in marked dose-dependent depression of body weight, hypoplastic changes in reproductive organs & accessory sex glands and hyperplasia of the forestomach mucosa. Reliability code 1. Reliable without restrictions.
	·
Remarks for Data Reliability References	The study was published in a peer-reviewed journal.  Herbert, C. D., Yuan, J. and Dieter, M.P. (1994). Comparison of the Toxicity of Cinnamaldehyde When Administered by Microencapsulation in Feed or by Corn Oil Gavage. Food and Chemical Toxicology 32(12): 1107-1115.

Substance Name	Cinnamaldehyde (98% pure)

**CAS No.** 104-55-2

**Method/guideline** A group of rats were fed a diet containing 0, 0.625, 1.25, 2.5,

5.0 or 10% Cinnamaldehyde microcapsules for 21 days.

**GLP** NG

**Year** 1994

Species/Strain B6C3F1 mice

**Sex** Male and Female

Route of administration Oral in feed

**Doses/concentration** A group of mice were fed a diet containing 0, 0.625, 1.25, 2.5,

5.0 or 10% Cinnamaldehyde microcapsules for 21 days.

**Exposure** 21 days

Frequency of treatment Continuous

**Remarks** A complete autopsy was performed on all animals that died,

and at the termination of the study on all treated and control animals. Body wt and the organ wt. of the liver, right kidney and

the spleen were determined.

NOAEL (NOEL) 1.25%

LOAEL (LOEL) 2.5%

Toxic response/effects by

dose level

Dose-related decrease in body wt, decrease in absolute liver and kidney wt., hyperplasia of the forestomach epithelium at

highest dose (10%) characterized by a focal thickening of the

stratified squamous epithelium, accompanied by

hyperkeratosis.

Statistical evaluations ANOVA

Conclusion Remarks Treatment of mice with microencapsulated cinnamaldehyde

resulted in dose-dependent depression of body weight and hyperplasia of the forestomach epithelium at highest dose (10%) characterized by a focal thickening of the stratified squamous epithelium, accompanied by hyperkeratosis.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was published in a peer-reviewed journal.

References Herbert, C. D., Yuan, J. and Dieter, M.P. (1994). Comparison of

the Toxicity of Cinnamaldehyde When Administered by Microencapsulation in Feed or by Corn Oil Gavage. Food and

Chemical Toxicology 32(12): 1107-1115.

Substance Name alpha-Hexylcinnamaldehyde

**CAS No.** 101-86-0

**Method/quideline** 90-day subchronic dermal toxicity study.

**GLP** NG

**Year** 1981

Species/Strain Sprague-Dawley Rats

**Sex** Male and Female

Route of administration Dermal

**Doses/concentration** 25 mg/kg

**Exposure** 90 days

Frequency of treatment Daily

Control Group and treatment Phenyl ethyl alcohol

**Remarks** 5% of the test substance in phenyl ethyl alcohol at a dose of 25

mg/kg was applied topically to the clipped backs of Sprague-Dawley rats (5 male and 5 female). A control group of 5 male and 5 female rats received phenyl ethyl alcohol (1 ml/kg). Body wt, hematology, clinical chemistry & urinalysis parameters were evaluated. All animals were examined grossly & liver & kidneys were weighed. Microscopic examination of the skin, liver,

kidney & spinal cord was conducted.

NOAEL (NOEL) 25 mg/kg

Toxic response/effects by

dose level

None

**Remarks for Results**One male rat died on day 14 with an evidence of a lung

infection. The death was not considered to be related to

treatment

Conclusion Remarks There was no evidence of toxicity induced by treatment with the

test articles.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**References** Moreno (1981a). 90 Day subacute dermal toxicity in rats with

hexyl cinnamic aldehyde, gamma-methyl ionone and phenyl

ethyl alcohol. Report to RIFM. Unpublished.

**Substance Name** p-t-Butyl-alpha-methylhydrocinnamaldehyde (97.8% pure),

liquid, colorless to pale yellowish.

**CAS No.** 80-54-6

Method/guideline Subchronic toxicity study. Test substance was administered to

albino rats by oral gavage. Six test groups consisting of 14 rats

per sex were dosed at 2, 5, 25 & 50 mg/kg once daily, 5

days/wk for 13 weeks.

**GLP** Yes

**Year** 1990

Species/Strain Rats, outbred

Sex Male and Female

Route of administration Oral gavage

**Doses/concentration** 0, 2, 5, 25 & 50 mg/kg/day,

**Exposure** 90 days

Frequency of treatment 5 days per week for 13 weeks

Control Group and treatment Rape oil 1 ml/kg

Post exposure observation

period Remarks 4 weeks

A satellite group was treated with 50 mg/kg and was observed during a post-treatment period of 4 weeks. Mortalities, general symptoms & body weights were recorded. Hematology & biochemistry determinations were performed. All rats were autopsied. Organs & tissues of the control rats & the rats treated w/50 mg/kg as well as liver of all rats, the testes & epididymides of all male rats & the adrenal glands of all female

rats were microscopically examined.

NOAEL (NOEL) 5.0 mg/kg

LOAEL (LOEL) 25 mg/kg

Toxic response/effects by

dose level

Statistical evaluations

Treatment related histopathology findings were spermatoceles

& testicular atrophy in male rats treated with 50 mg/kg

Dunn-test, Jonck-heere-test, U-test

Remarks for Results Deaths related to treatment did not occur throughout the test

and follow-up period. Loss of hair was seen in female rats treated with 50 mg/kg. The body wt development of tats of all test groups took a normal course throughout the test and follow-up period. The treatment did not change hematological parameters. In male and female rats treated with 25 and 50 mg/kg, the plasma cholinesterase was reversibly decreased and the plasma cholesterol levels were lower than in control

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Conclusion Remarks Treatment with test material resulted in spermatoceles and

testicular atrophy in male rats at the dose of 50 mg/kg. Also, a decrease in the plasma cholinesterase activity and plasma cholesterol was seen in rats treated with the test material at the

dose of 25 mg/kg and above.

**Data Qualities Reliabilities** Reliability code 1. Reliable without restrictions.

Remarks for Data Reliability The study was conducted in accordance with EPA Guidelines,

OECD Guidelines and Swiss Guidelines.

**References** Givaudan-Roure (1990d) A supplementary study with *p*-t-butyl-

alpha-methylhydrocinnamic aldehyde on rats for determining acetylcholinesterase and cholinesterase activity of blood plasma, erythrocytes, liver and brain tissue. Unpublished

Report to RIFM.

**Substance Name** p-t-Butyl-alpha-methyldihydrocinnamaldehyde

**CAS No.** 80-54-6

**Method/guideline** Groups of 8 male rats were treated with 25, 50, 100, 200 & 400

mg/kg/day of the test substance orally for 5 consecutive days.

**GLP** Likely

Year 1990

Species/Strain Rats

Sex Male

Route of administration Oral (gavage)

Doses/concentration 0, 25, 50, 100, 200 & 400 mg/kg/day

**Exposure** 5 consecutive days

Frequency of treatment Daily

**Control Group and treatment** Yes

Remarks This is a follow-up study on the previous study by the same

group with similar results

Toxic response/effects by

**Remarks for Results** 

dose level mg/kg & above, morphological alterations in the seminiferous

epithelium preceded the formation of detectable spermatoceles. No deaths reported, other observations reported: shaggy fur, hunched posture, hematuria, paresis of the forelegs, initial disturbance of weight development @ 50, 100 & 200 mg/kg/day which recovered on day 4; continued loss of body weight @ 400 mg/kg; At autopsy, delineation of hepatic lobules, small prostate and seminal vesicles, and reddened testes were seen. Testes weight was decreased in rats treated with 100 mg/kg

Disturbed the spermatogenesis and spermiogenesis @ 100

and above, histological examination of the testes revealed injuries of seminiferous epithelium that means degeneration and loss of germ cells in rats treated with 50 mg/kg and above.

**Conclusion Remarks** Administration of the test substance for 5 consecutive days

resulted in disturbance of the spermatogenesis and spermiogenesis. The, morphological alterations in the seminiferous epithelium preceded the formation of detectable

spermatoceles.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability Two of the four control rats displayed a disturbance of

spermiogenesis with desquamation of young spermatids. Also,

authors states that rat seems to be much more prone to

spermatoceles than e.g. the mouse, therefore the rat might be a bad model for detecting epididymal side effects of chemicals. Authors also claim that the rat was found to be only species to suffer from adverse testicular and epididymal effect from

exposure to the test chemical.

References Givaudan Roure (1990c). Re-evaluation of testicular and

epididymal side effects caused by p-t-butyl alpha-

methyldihydrocinnamic aldehyde in rats following short (5 days) and subchronic (13 weeks) oral administration. Unpublished.

Report to RIFM.

p-t-Butyl-alpha-methyldihydrocinnamaldehyde Substance Name

**CAS No.** 80-54-6

Method/guideline 13-weeks Subchronic study. Groups of 14 male and 14 female

rats were treated by oral gavage with 0, 5, 25, and 50 mg/kg for

five days per week for 13 consecutive weeks.

GLP Likely

**Year** 1990

Species/Strain Rat

Sex Male

Route of administration Oral (gavage)

**Doses/concentration** 0, 2, 5, 25 and 50 mg/kg/day

Exposure 13 weeks

Frequency of treatment 5 days per week for 13 weeks

**Control Group and treatment** Yes

Post exposure observation

period Remarks 4 weeks

The rats were sacrificed with exception of 4 control rats per sex and a satellite group of 14 rats per sex treated with 50 mg/kg. These rats were necropsied after a treatment-free period of

approximately 4 weeks.

NOAEL (NOEL) 5 mg/kg/day

LOAEL (LOEL) 25 mg/kg/day

Toxic response/effects by

dose level

Necropsy findings comprised spermatoceles and the

occurrence of small testes in male rats treated with 50 mg/kg. Treatment-related histopathology findings were spermatoceles

and testicular atrophy in male rats treated with 50 mg/kg.

Remarks for Results No treatment related deaths, Other treatment related

observation included: Loss of hair in female rats, reversible decrease in cholinesterase activity and the plasma cholesterol levels in male and female rats, Absolute and relative weights were elevated in male and female rats treated with 25 and 50 mg/kg. The absolute and relative weights of adrenal glands were elevated in female rats treated with 25 and 50 mg/kg. Administration of the test substance for 5 consecutive days

Conclusion Remarks Administration of the test substance for 5 consecutive days resulted in disturbance of the spermatogenesis and

spermiogenesis. The morphological alterations in the seminiferous epithelium preceded the formation of detectable

spermatoceles.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** Two of the four control rats displayed a disturbance of

spermiogenesis with desquamation of young spermatids. Also, authors states that rat seems to be much more prone to spermatoceles than e.g. the mouse, therefore the rat might be a

bad model for detecting epididymal side effects of chemicals. Authors also claim that the rat was found to be only species to suffer from adverse testicular and epididymal effect from

exposure to the test chemical.

**References** Givaudan Roure (1990c). Re-evaluation of testicular and

epididymal side effects caused by p-t-butyl alpha-

methylhydrocinnamic aldehyde in rats following short (5 days) and subchronic (13 weeks) oral administration. Unpublished.

Report to RIFM.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	Test substance was administered orally to white rats for 25 day.

GLP NG

**Year** 1974

Species/Strain White rat

Sex Male

Route of administration Oral (gavage)

**Doses/concentration** 0.02 LD50 (LD50 = 3400 mg/kg)

**Exposure** 25days

Control Group and treatment Sunflower seed oil

**Remarks** Following parameters were monitored: Plasma Cholinesterase

activity, serum aldolases activity, sorbitol dehydrogenase, aspartate and alanine aminotransferase, content of SH groups,

total protein level in the blood serum.

Toxic response/effects by

dose level

No effects were reported.

**Conclusion Remarks** 

No adverse effects were seen in the rats treated with cinnamaldehyde for 25 days at a dose of 0.02LD50.

Data Qualities Reliabilities Reliability cod

Reliability code 3. Data not reliable.

Remarks for Data Reliability

Original article in Russian. Very few details given.

References

Zaitsev, A.N. and Rakhmanina, N.L. (1974). Some Data on the Toxic Properties of Phenylethyl and Cinnamyl Alcohols. Vopr

Pitaniya 6: 48-53.

Substance Name	alpha-Hexylcinnamaldehyde
CAS No.	101-86-0
Method/guideline	Test material was applied percutaneously to the shaved dorsa of 10 male Sprague-Dawley rats at dose levels of 0.15, 0.375, 0.75, 1.5 and 3.0 g/kg/day for 28 consecutive days,

**GLP** Yes

**Year** 1980

Species/Strain Sprague-Dawley rats

**Sex** Male

Route of administration Percutaneous

**Doses/concentration** 0.15, 0.375, 0.75, 1.5 and 3.0 g/kg/day

**Exposure** 28 days

Frequency of treatment Daily

Control Group and treatment None

Post exposure observation

period Remarks None

Parameter monitored: body wt., food consumption, hematology, blood chemistry (BUN), serum glutamic pyruvic transaminase, serum glutanic oxalacetic transaminase, total bilirubin, fasting serum glucose, serum alkaline phosphatase), gross pathology,

histopathology.

LOAEL (LOEL)

0.15 g/kg/kg

Toxic response/effects by

dose level

Erythema and eschar formation with cracking and dryness @all doses, hyperirritability @ all doses except 0.375 g/kg/day, reduced body wt @ 1.5 & 3.0 g/kg/day, depressed food intake @ 3.0 g/kg/day, dose-related negative effect on clotting time & white blood cell count, shift in the proportion of segmented neutrophils to lymphocytes @ 1.5 & 3.0 g/kg, increase in BUN, SAP, SGPT, SGOT & decrease in Glucose, thickening of the skin & erythema of dermis & epidermis, body emaciation, congested lungs, GI irritation, decrease in absolute & relative thymus & spleen, dermatitis with mild to severe hyperkeratosis at all doses except 0.15 g/kg, focal dilation of tubules in kidney

@~0.75~&~1.5~g/kg, sub-acute to chronic necrotizing &~

hemorragic enteritis

**Statistical evaluations** No statistical evaluation was done.

**Remarks for Results**Because small number of animals (2 per group) no statistical

evaluation was done.

**Conclusion Remarks** Repeated percutaneous administration of *alpha* hexylcinnamic

aldehyde resulted in changes in gross pathology,

histopathology, clinical and biochemical chemistry and

hematological parameters.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

Remarks for Data Reliability The study was conducted in accordance with GLP.

**References** Lough, R., Owston, E., Bier, C., and Qureshi, S. (1980). A

Range finding evaluation of the toxicity of Hexyl Cinnamic aldehyde Administered percutaneously in the rat. Unpublished.

Bio-Research Laboratories Ltd. Report to RIFM.

Substance Name	alpha-Amylcinnamaldehyde
CACNO	•
CAS No.	122-40-7
Method/guideline GLP	15 Male & 15 Female rats of the FDRL strains were fed diet containing 2% test substance diluted in cotton-seed oil for 12 weeks. At 90 days, autopsy was performed. Hematological and blood chemical determinations were also made.  Pre GLP
Year	1965
Species/Strain	FDRL Strain Rats
Sex	Male and Female
Route of administration	Feed
Doses/concentration	2% in feed
Exposure	12 weeks
Frequency of treatment	Feed diet with test material for 12 weeks
Control Group and treatment	Feed without tests material
Post exposure observation period	NG
NOAEL (NOEL)	2%
Statistical evaluations	NG
Remarks for Results	No treatment related adverse effects were noted in the
<b>Conclusion Remarks</b>	parameters measured. This study demonstrates a NOAEL in rats was shown to be 2% in feed.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.
References	Oser, B.L., Carson, S, and Oser, M. (1965) Toxicological Tests on Flavouring Matters. Fd. Cosmet. Toxicol. 3: 563-569.

## 4.5 Reproductive Toxicity

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline	NG
Test Type	Two generations

**GLP** NG

**Year** 1965

Species/Strain Not given

Sex Female

Route of administration Unreported

**Duration of test** 223 & 210 days

**Doses/concentration** 2 mg cinnamaldehyde every other day

**Premating Exposure period** 

for males

NA

Frequency of treatment Continuous

Control Group and treatment Not mentioned

**Remarks for Test Conditions** Article in Romanian. English abstract contains very few details.

Parameters monitored: body weight, reproduction ability (no. of pregnant females, no.& weight of the young one at birth), the development & viability of the young animals, the protein & lipid

contents of liver & liver activity.

Remarks for Results Treatment resulted in significant (p<0.01) 20-22% increase in

the lipid content of the liver as compared to control groups. The other indicators were not affected. No details were given whether the observed effect was in offspring or Parents. Article

in Romanian.

Conclusion Remarks Administration of the test substance caused in increase in liver

lipid content in the unspecified group.

**Data Qualities Reliabilities** Reliability code 3. Data not reliable.

Remarks for Data Reliability Article in Romanian. The English abstract contains very few

details.

**General Remarks** Article is in Romanian. Need English translation of more details.

**References** Sporn A. (1965). Investigation of the Toxicity of Cynamic

Aldehyd. Igiena 14(6): 339-346.

Substance Name	Cinnamaldehyde
CAS No.	104-55-2

Method/guideline Cinnamic aldehyde was administered by gavage to Sprague-

Dawley rats on days 7-17 of pregnancy at doses of 5, 25 or 250

mg/kg/day.

**Test Type** Pre-Natal (Segment II) Toxicity Study.

**GLP** NG

**Year** 1989

Species/Strain Sprague-Dawley rats

**Sex** Female

Route of administration Gavage

**Duration of test** Days 7-17 of pregnancy

**Doses/concentration** 0, 5, 25 or 250 mg/kg/day

**Premating Exposure period** 

for males

None

**Premating Exposure period** 

for females

None

Frequency of treatment Daily

Control Group and treatment Vehicle (olive oil)

NOAEL (NOEL) None

LOAEL (LOEL) 5 mg/kg/day

Actual dose received by dose level and sex Parental data and F1

0, 5, 25 or 250 mg/kg/day

No signs of maternal toxicity, decreased weight gain between

day 7 & 20 with decrease in food intake.

Offspring toxicity F1 and F2 Increased incidence of poor cranial ossification, decreased

ossification of tympanic bulla at 25 or 250 mg/kg/day, increased incidence of dilated pelvis/reduced papilla in kidney, increased incidence of reduced cranial ossification, dilated ureter. One

case of facial malformation & few cases of

hypoplastic/dysplastic kidney.

Statistical evaluations Kruskal-Wallis test, Mann-Wittney test

Remarks for Results Authors abstract state "significant increases of the incidences of

dilated pelvis/reduced papilla in the kidney, dilated ureters>2 abnormal sternebrae per fetus were detected in the 2-mg/kg group." However no such dose group (2-mg/kg) is reported in

either the methodology or the Results section.

**Conclusion Remarks** Administration of Cinnamaldehyde to pregnant rats resulted in

increased incidence of poor cranial ossification and reduced ossification of the tympanic bulla. Significant increases of the incidences of dilated pelvis/reduced papilla in the kidney, ureters > 2 abnormal sternebrae per fetus were also reported.

**Data Qualities Reliabilities** Reliability code 2. Reliable with restrictions.

**Remarks for Data Reliability** The study was published in a peer-review journal.

**General Remarks** The changes in treated groups might have been influenced by

the greater litter size in the higher dose groups (There was significantly higher pre-implantation loss in control as compared

to the treated groups).

References Mantovani, A., Stazi, A.V., Macri, C., Ricciardi, C., Piccioni, A.

and Badellino, W. (1989). Pre-Natal (Segment II) Toxicity Study of Cinnamic Aldehyde in the Sprague-Dawley Rats. Food and

Chemical Toxicology 27(12): 781-786.

Substance Name	Instead of Cinnamaldehyde, structurally related chemicals, Cinnamic alcohol and cinnamic acid were used in this study
CAS No.	104-55-2
Method/guideline Test Type	The chemicals were studied at doses of 0.02 and 0.002 LD50 value; 53.5 mg/kg cinnamic alcohol and 50 mg/kg cinnamic acid. The animals were exposed to the test chemical during the entire pregnancy.  Reproductive toxicity.
GLP	NG
Year	1975
Species/Strain	Albino rat
Sex	Female
Route of administration	Oral
Duration of test	20 days
Doses/concentration	53.5 mg/kg cinnamic alcohol and 50 mg/kg cinnamic acid
Premating Exposure period for males	None
Premating Exposure period for females	None
Frequency of treatment	Daily
Control Group and treatment	Yes
Remarks for Test Conditions  Data Qualities Reliabilities	There were 14-15 female rats in each group. On the 20th day of pregnancy 6-9 rats from each group were decapitated, the embryos were taken from the uterus and studied. The remaining pregnant rats were left until the natural birth and the development of the progeny was observed during the postnatal period for one month. The parameters monitored: embryonic mortality, number of live embryos, birth weight, length, number of external and internal anomalies in the development of the embryos.  Reliability code 2. Reliable with restrictions.
Offspring toxicity F1 and F2	Cinnamic alcohol and cinnamic acid administration perorally to
Conclusion Remarks References	rats during the whole pregnancy is doses of 53.5 and 50 mg/kg caused no embryotoxic effect.  Cinnamic alcohol and cinnamic acid administration perorally to rats during the whole pregnancy is doses of 53.5 and 50 mg/kg caused no embryotoxic effect.  Zaitsev, A. N. and Maganova, N. B. (1975). Embryotoxic Action of Some Food Aromatizers. Voprosy Pitaniya 3: 6468.

## 4.6 Developmental Toxicity

Substance Name	Cinnamaldehyde
CAS No.	104-55-2
Method/guideline Test Type	The study was conducted in two phases: initial dose-finding study followed by a reproductive phase, which employed a single dose level. In both phases treatment was administered by gavage using a standard dosing volume of 10 ml/kg. Developmental Toxicity Test
GLP	NG
Year	1987
Species/Strain	CD1 mice
Sex	Female
Route of administration	Oral (gavage)
<b>Duration of test</b>	From Gestation Day 6-13
Doses/concentration	1200 mg/kg/day
Premating Exposure period for males	None
Premating Exposure period for females	None
Frequency of treatment	Daily
Control Group and treatment	Corn oil
Remarks for Test Conditions	For Phase I, test chemical was tested at five dose levels using ten virgin female mice for 8 consecutive days. For the Reproductive phase, the LD10 predicted on the basis of dose-finding results was the single dose used. Treatment in the reproductive phase were administered once daily on Gestation day 6-13
NOAEL (NOEL)	1200 mg/kg/day
Parental data and F1	As compared to controls, no changes were seen in: Number of dead/total; % body weight change and delivery of viable litter.
Offspring toxicity F1 and F2	As compared to control, no changes were seen in: Number of stillborn/litter; %survival; birth weight and weight gain.
Statistical evaluations	2-tail ANOVA, 2-tail Fischer's exact test,
Conclusion Remarks	Administration of Cinnamaldehyde to pregnant female mice (gestation day 6-13) did not produce any maternal, fetal or neonatal toxicity.
Data Qualities Reliabilities	Reliability code 2. Reliable with restrictions.
Remarks for Data Reliability	The study was published in a peer-reviewed journal.

## References

Hardin, B.D.m Schufer, R.L., Burg, J. R., Booth, G.M., Hazelden, K.P., MacKenzie, K.M., Piccirillo, V. J. and Smith, K.N. (1987). Evaluation of 60 Chemicals in a Preliminary Developmental Toxicity Test. Teratogenesis, Carcinogenesis and Mutagenesis 7: 29-48.